(FILE 'HOME' ENTERED AT 10:07:38 ON 12 SEP 2003)

FILE 'REGISTRY' ENTERED AT 10:07:50 ON 12 SEP 2003 11 S METFORMIN OR GLIPIZIDE

FILE 'HCAPLUS, MEDLINE, BIOSIS, EMBASE, JAPIO, USPATFULL' ENTERED AT 10:08:52 ON 12 SEP 2003 4879 S 338752-31-1/RN OR 338752-30-0/RN OR 88159-36-8/RN OR 29

4879 S 338752-31-1/RN OR 338752-30-0/RN OR 88159-36-8/RN OR 29094-61 1713 S 1115-70-4/RN OR MET FORMIN OR GLUFORMIN OR GLYFORMIN OR GLUME 1717 S L3 OR 58840-24-7/RN OR 53950-18-8/RN OR 38950-16-2/RN OR 344

L5 404 S L4 AND L2

Ll

L2 L3 L4

L6

394 DUP REM L5 (10 DUPLICATES REMOVED)

L7 183 S L6 AND (SINGLE DOSAGE OR TABLE OR CAPSULE OR SINGLE DOSE)

L8 183 FOCUS L7 1-

6 303 146

11 ANSWER 1 OF 56 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:944287 HCAPLUS

DOCUMENT NUMBER: 138:100731

Glyburide/metformin combination TITLE:

product is safe and efficacious in patients with type

2 diabetes failing sulfonylurea therapy

Blonde, L.; Rosenstock, J.; Mooradian, A. D.; Piper, AUTHOR (S):

B.-A.; Henry, D.

Ochsner Clinic Foundation, New Orleans, LA, USA CORPORATE SOURCE:

SOURCE:

Diabetes, Obesity and Metabolism (2002), 4(6), 368-375

CODEN: DOMEF6; ISSN: 1462-8902

PUBLISHER: Blackwell Science Ltd.

Journal DOCUMENT TYPE: LANGUAGE: English

The aim was to compare the efficacy, safety and tolerability of a fixed combination glyburide/metformin prepn. with those of qlyburide or metformin alone in patients with type 2 diabetes inadequately controlled by sulfonylurea, diet and exercise. In this 16-wk, randomized, double-blind, parallel group study, 639 patients with inadequate glycemic control on at least half-maximal dose of sulfonylurea were randomly assigned to: glyburide 10 mg b.i.d. (n = 164); metformin 500 mg (n = 153); glyburide/metformin 2.5 mg/500 mg (n = 160); or glyburide/metformin 5 mg/500 mg (n = 162). Titrn. was allowed to max. doses of 2000 mg for metformin or 10 mg/2000 mg and 20 mg/2000 mg for glyburide/metformin 2.5 mg/500 mg and 5mg/500 mg resp. The primary outcome measure was HbA1c level after 16 wk; secondary end-points included fasting and 2-h post-prandial plasma glucose. Adverse events (AEs) were recorded and summarized by treatment group. Both strengths of glyburide/metformin equally reduced mean HbAlc by 1.7% more than did glyburide alone (p < 0.001), and by 1.9% more than did metformin alone (p < 0.001). Final mean fasting plasma glucose concns. were also lower in both glyburide/ metformin groups than in the glyburide (-2.8 mmol/l, -51.3 mg/dL; p < 0.001) and metformin groups (-3.6 mmol/l, -64.2 mg/dL; p < 0.001). Safety and tolerability were similar across all treatment groups, except for a higher incidence of gastrointestinal AEs in the metformin monotherapy group, and more patients reporting mild or moderate symptoms of hypoglycemia while taking glyburide/metformin. Both glyburide/metformin tablet strengths produced, with equal efficacy, significantly better glycemic control than monotherapy with either agent. These data also confirm that glycemic efficacy does not require maximal sulfonylurea doses in combination with metformin.

THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 20 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT CCESSION NUMBER:

2002:499506 HCAPLUS

DOCUMENT NUMBER:

137:119415

TITLE:

Simultaneous glyburide/metformin

therapy is superior to component monotherapy as an initial pharmacological treatment for type 2 diabetes Garber, A. J.; Larsen, J.; Schneider, S. H.; Piper, B.

AUTHOR (S):

A.; Henry, D.

CORPORATE SOURCE:

Baylor College of Medicine and The Methodist Hospital,

Houston, TX, 77030, USA

SOURCE:

Diabetes, Obesity and Metabolism (2002), 4(3), 201-208

CODEN: DOMEF6; ISSN: 1462-8902

PUBLISHER:

Blackwell Science Ltd.

DOCUMENT TYPE: LANGUAGE:

Journal English

The aim of this study was to evaluate whether simultaneous initial treatment of both insulin resistance and impaired .beta.-cell insulin secretion with glyburide/metformin tablets is superior to monotherapy with each component agent. In this randomized, parallel-group, placebo-controlled, multicenter study, 806 patients with type 2 diabetes (mean duration, 3 yr) who had failed diet and exercise were randomly assigned to 4 wk of therapy with placebo, glyburide 2.5 mg, metformin 500 mg, glyburide/metformin 1.25/250 mg, or glyburide/metformin 2.5/500 mg once daily. Doses were then titrated over 8 wk based on glycemic response. The primary outcome measure was change from baseline in mean HbA1c after 20 wk. Changes in fasting plasma glucose, lipids and body wt. were also assessed along with 2-h postprandial glucose and insulin values after a standardized meal. At week 20, patients taking glyburide/metformin 1.25/250 mg or 2.5/500 mg tablets had greater redns. in HbAlc levels (-1.48% and -1.53% resp.) compared with placebo (-0.21%; both p < 0.001), glyburide (-1.24%; p = 0.016 and p = 0.004 resp.) or metformin (-1.03%; both p < 0.016)0.001). Fasting plasma glucose concns. were reduced more in both glyburide/metformin groups compared with placebo and metformin (p < 0.001); patients in both combination therapy groups also had significantly lower postprandial glucose concns. compared with placebo, glyburide and metformin. Initial combination treatment with glyburide/metformin tablets produces greater improvements in glycemic control than either glyburide or metformin monotherapy. The superiority of initial therapy with glyburide/

metformin tablets may arise from simultaneous treatment of both

8

REFERENCE COUNT:

pathophysiol. defects of type 2 diabetes. THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT L11 ANSWER 3 OF 56 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:842385 HCAPLUS

DOCUMENT NUMBER: 137:332944

TITLE: Lipid effects of glyburide/metformin

tablets in patients with type 2 diabetes mellitus with

poor glycemic control and dyslipidemia in an

open-label extension study

AUTHOR(S): Dailey, George E., III; Mohideen, Pharis; Fiedorek,

Fred T.

CORPORATE SOURCE: Diabetes and Endocrinology, Scripps Clinic, La Jolla,

CA, USA

SOURCE: Clinical Therapeutics (2002), 24(9), 1426-1438

CODEN: CLTHDG; ISSN: 0149-2918

PUBLISHER: Excerpta Medica, Inc.

DOCUMENT TYPE: Journal LANGUAGE: English

Because both type 2 diabetes and elevated plasma lipid levels are important independent risk factors for cardiovascular disease and coronary heart disease, the choice of an antihyperglycemic agent for patients with type 2 diabetes-in whom abnormal plasma lipid levels are often seen-should take into account effects on lipids as well as on markers of glycemic control. This study assessed the effects on lipid levels of glyburide/metformin tablets in the treatment of type 2 diabetes, particularly in a group of patients who had poor glycemic control and dyslipidemia at baseline. This 52-wk, open-label study was an extension of a 32-wk, double-blind, placebo-controlled study. The patient population was drawn from 3 groups: those who completed the double-blind study, those who were discontinued from the double-blind study, and those who were ineligible for the double-blind study based on predefined measures of glycemic control (screening fasting plasma glucose >240 mg/dL and glycosylated Hb [HbAlc] .ltoreq.12%, or HbAlc 11%-12%) and were directly enrolled in the open-label extension study. Patients with an HbAlc of <9% received glyburide/metformin tablets 1.25 mg/250 mg BID; those with an HbA1c .gtoreq.9% received glyburide /metformin tablets 2.5 mg/500 mg BID. Changes in total cholesterol (TC), low-d. lipoprotein cholesterol (LDL-C), high-d. lipoprotein cholesterol (HDL-C), and triglyceride (TG) levels were assessed for 52 wk. The study population included 828 patients: 515 who completed the double-blind study, 138 who were discontinued from the double-blind study, and 175 who were enrolled directly. Direct enrollees had poor glycemic control and dyslipidemia at baseline. Improvements in plasma lipid levels were seen as early as week 13. At week 52, the mean change in TC from baseline was -8.0 mg/dL for the total population (95% CI, -10.9 to -5.2; P < 0.05) and -23.2 mg/dL for direct enrollees (95% CI, -30.1 to -16.4; P < 0.05). The mean decrease in LDL-C from baseline for the total population was 2.86 mg/dL (95% CI, -5.3 to -0.4; P < 0.05), compared with a redn. of 13.3 mg/dL for direct enrollees (95% CI, -18.5 to -8.1; P < 0.05). Mean HDL-C levels were minimally affected. Mean TG levels decreased by 27.8 mg/dL for the entire population (95% CI, -4.2.9to -12.8; P < 0.05) and by 99.7 mg/dL for direct enrollees (95% CI,-152.5 to-46.8; P < 0.05). In this open-label extension study, treatment with glyburide/metformin tablets for type 2 diabetes had a durable, favorable effect on lipid levels, particularly in those with poor glycemic control and dyslipidemia at baseline.

REFERENCE COUNT: 39 THERE ARE 39 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ACCESSION NUMBER:

2003:557424 HCAPLUS

TITLE:

Simultaneous glyburide/metformin

therapy is superior to component monotherapy as an

initial pharmacological treatment for type 2 diabetes.

[Erratum to document cited in CA137:119415]

Garber, A. J.; Larsen, J.; Schneider, S. H.; Piper, B.

A.; Henry, D.

CORPORATE SOURCE:

Baylor College of Medicine and The Methodist Hospital,

Houston, TX, 77030, USA

SOURCE:

Diabetes, Obesity and Metabolism (2002), 4(4), 286

CODEN: DOMEF6; ISSN: 1462-8902

PUBLISHER:

AUTHOR (S):

Blackwell Science Ltd.

DOCUMENT TYPE:

Journal; Errata

LANGUAGE:

English

An erratum.

L11 ANSWER 5 OF 56 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:842383 HCAPLUS

DOCUMENT NUMBER: 137:332943

TITLE: Durability of efficacy and long-term safety profile of

glyburide/metformin tablets in

patients with type 2 diabetes mellitus: an open-label

extension study

AUTHOR(S): Garber, Alan J.; Bruce, Simon; Fiedorek, Fred T.

CORPORATE SOURCE: Baylor College of Medicine and the Methodist Hospital,

Houston, TX, USA

SOURCE: Clinical Therapeutics (2002), 24(9), 1401-1413

CODEN: CLTHDG; ISSN: 0149-2918

PUBLISHER: Excerpta Medica, Inc.

DOCUMENT TYPE: Journal LANGUAGE: English

Intensive qlycemic control substantially reduces the microvascular and macrovascular complications of type 2 diabetes mellitus, although less than half of patients with diabetes achieve the target glycosylated Hb (HbA1c) value recommended by the American Diabetes Assocn. Because monotherapy with an oral agent does not address the multiple pathophysiol. defects of diabetes, use of combination therapy appears to be warranted. A previous 32-wk, randomized, double-blind, placebo-controlled trial found that treatment with glyburide/metformin tablets was assocd. with greater redns. in HbAlc values compared with glyburide monotherapy, metformin monotherapy, and placebo. This study evaluated the durability of efficacy and long-term safety profile of therapy with glyburide/metformin tablets over 52 wk. Patients enrolled in this open-label extension study were drawn from 3 groups: those who completed the 32-wk double-blind study, those who were discontinued from the double-blind study, and those who were ineligible for the double-blind study and were enrolled directly in the open-label extension study. Patients with an HbA1c of <9% received glyburide /metformin 1.25 mg/250 mg tablets BID, and those with an HbA1c of .gtoreq.9% received glyburide/metformin 2.5 mg/500 mg tablets BID. Primary efficacy variables included changes from baseline in HbAlc, fasting plasma glucose (FPG), and body wt. at week 52. Safety was assessed based on adverse-event data and the results of phys. examns. and lab. tests. A total of 828 patients were enrolled in the study: 515 who completed the 32-wk double-blind study, 138 who were discontinued from the double-blind study, and 175 who were directly enrolled. At week 52, the mean HbAlc value for the entire population had decreased from a baseline value of 8.73% to 7.04% (95% CI, -1.81 to -1.58). Patients who were enrolled directly had the poorest glycemic control at baseline and experienced the greatest redn. in HbA1c (-3.35%; 95% CI, -3.61 to -3.10). A redn. in mean FPG for the total population was obsd. as early as week 2, from 201 to 141 mg/dL (95% CI, -63.0 to -55.7). Symptoms of hypoglycemia occurred in 19.9% (165/828) of patients, although only one third of these patients had a documented finger-stick blood glucose value of .ltoreg.50 mg/dL. In this 52-wk, open-label extension study, glyburide/ metformin tablets were well tolerated and effective in patients with type 2 diabetes. They provided rapid and sustainable redns. in HbAlc values and FPG concns.

REFERENCE COUNT: 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CCESSION NUMBER: 2003:657740 HCAPLUS

TITLE: Efficacy of glyburide/metformin

tablets compared with initial monotherapy in type 2

diabetes

AUTHOR(S): Garber, Alan J.; Donovan, Daniel S., Jr.; Dandona,

Paresh; Bruce, Simon; Park, Jong-Soon

CORPORATE SOURCE: Baylor College of Medicine and the Methodist Hospital,

Houston, TX, 77030, USA

SOURCE: Journal of Clinical Endocrinology and Metabolism

(2003), 88(8), 3598-3604

CODEN: JCEMAZ; ISSN: 0021-972X

PUBLISHER: Endocrine Society

DOCUMENT TYPE: Journal LANGUAGE: English

Many patients with type 2 diabetes fail to achieve or maintain the American Diabetes Assocn.'s recommended treatment goal of glycosylated Hb levels. This multicenter, double-blind trial enrolled patients with type 2 diabetes who had inadequate glycemic control [glycosylated Hb A1C (A1C), >7% and <12%) with diet and exercise alone to compare the benefits of initial therapy with glyburide/metformin tablets vs. metformin or glyburide monotherapy. Patients (n = 486) were randomized to receive glyburide/metformin tablets (1.25/250 mg), metformin (500 mg), or glyburide (2.5 mg). Changes in A1C, fasting plasma glucose, fructosamine, serum lipids, body wt., and 2-h postprandial glucose after a standardized meal were assessed after 16 wk of treatment. Glyburide/metformin tablets caused a superior mean redn. in A1C from baseline (-2.27%) vs. metformin (-1.53%) and glyburide (-1.90%) monotherapy (P = 0.0003). Glyburide/metformin also significantly reduced fasting plasma glucose and 2-h postprandial glucose values compared with either monotherapy. The final mean doses of glyburide/metformin (3.7/735 mg) were lower than those of metformin (1796 mg) and glyburide (7.6 mg). First-line treatment with glyburide/metformin tablets provided superior glycemic control over component monotherapy, allowing more patients to achieve American Diabetes Assocn. treatment goals with lower component doses in drug-naive patients with type 2 diabetes.

REFERENCE COUNT: 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ACCESSION NUMBER:

2003:564950 HCAPLUS

TITLE:

Glyburide/metformin tablets: a new

therapeutic option for the management of Type 2

diabetes

AUTHOR(S):

Dailey, George E.

CORPORATE SOURCE:

10666 N. Torrey Pines Road, La Jolla, CA, 92037, USA

SOURCE:

Expert Opinion on Pharmacotherapy (2003), 4(8),

1417-1430

CODEN: EOPHF7; ISSN: 1465-6566

Ashley Publications Ltd.

DOCUMENT TYPE:

PUBLISHER:

Journal English

LANGUAGE:

Oral antidiabetic combination therapy is a proven means of establishing glycemic control in the hyperglycemic, Type 2 diabetic patient, but co-administering two oral antidiabetic agents sep. may hinder compliance with therapy. A new single-tablet of glyburide/

metformin combination therapy (Glucovance, Bristol-Myers Squibb, Inc.) has recently been developed, which addresses the primary defects of Type 2 diabetes: .beta.-cell dysfunction and insulin resistance. The

glyburide/metformin tablet, taken with meals, is designed to optimize the absorption of glyburide and to address the postprandial glucose rise. Glyburide/metformin

tablets are more effective in controlling fasting and postprandial glycemia than its component monotherapies, at lower doses of metformin and glyburide compared with monotherapy because of the synergy between its glyburide and metformin components. Moreover, a double-blind study showed that glyburide/metformin tablets are more effective

than a free combination of glyburide co-administered with metformin in controlling postprandial glucose. Retrospective analyses suggested that glyburide/metformin tablets control glycated Hb (A1C)

more effectively than a free combination of glyburide co-administered with metformin, at lower mean doses of glyburide and metformin. The incidence of side effects is lower than sep. component therapy for any given AlC. Glyburide/metformin tablets are an effective option for optimizing the control of blood glucose in Type 2 diabetic patients and appear to enhance adherence to therapy.

ACCESSION NUMBER: 2003:51312 HCAPLUS

DOCUMENT NUMBER: 138:117496

TITLE: Pharmacokinetics and pharmacodynamics of

glyburide/metformin tablets

(Glucovance) versus equivalent doses of glyburide and

metformin in patients with type 2 diabetes

AUTHOR(S): Donahue, Stephen R.; Turner, Kenneth C.; Patel,

Shardul

CORPORATE SOURCE: Department of Clinical Discovery, Bristol-Myers Squibb

Pharmaceutical Research Institute, Princeton, NJ, USA Clinical Pharmacokinetics (2002), 41(15), 1301-1309

SOURCE: Clinical Pharmacokinetics (2002), 41(1 CODEN: CPKNDH; ISSN: 0312-5963

PUBLISHER: Adis International Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

Objective: To compare the effects of two different formulations of glibenclamide (glyburide) combined with metformin on postprandial glucose excursions, and to assess their pharmacokinetics. The formulations were a combination glibenclamide/metformin tablet (Glucovance; controlled-particle-size glibenclamide and metformin) vs. glibenclamide (Micronase) and metformin (Glucophage) coadministered sep. Design: A randomized, double-blind, two-way crossover study in which patients with type 2 diabetes received either glibenclamide/metformin 2.5/500mg tablets or glibenclamide 2.5mg with metformin 500mg twice daily for 14 days. After a 2-wk washout, patients were crossed over to the other treatment for 14 days. Patients consumed standardized meals on the days when pharmacokinetic and pharmacodynamic evaluations were performed. Participants: Forty patients with type 2 diabetes were enrolled; 37 were randomized (18 men, 19 women) and 35 completed the study. Mean age was 58 yr; mean body mass index was 31 kg/m2. The baseline glycated Hb (HbA1c) was 9.3% for both treatment groups. Main outcome measure: Two-hour postprandial glucose excursion (PPGE) was used to assess postprandial glucose dynamics. Results: Treatment with glibenclamide/metformin resulted in a significantly smaller mean PPGE than was attained by treatment with glibenclamide plus metformin, according to measurements taken after the day 14 afternoon standardized meal (89.5 vs. 117.4 mg/dL, p = 0.011). The mean glibenclamide peak concn. (Cmax) was significantly greater (.apprx.16%) after glibenclamide/metformin treatment on both days 1 and 14. Glibenclamide/metformin treatment was assocd. with a 2-fold greater area under the concn.-time curve to 3 h for glibenclamide (AUC3) [p < 0.001], although the AUC over the administration interval was equiv. for both formulations. Conclusion: In patients with type 2 diabetes, glibenclamide/metformin resulted in lower PPGE, suggesting that the higher glibenclamide AUC3 obsd. with this formulation may contribute to better postprandial glycemic control than is attained by glibenclamide plus metformin sep.

REFERENCE COUNT: 16 THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

MEDLINE on STN L11 ANSWER 14 OF 56 ACCESSION NUMBER: 2003077460 MEDLINE

PubMed ID: 12589230 DOCUMENT NUMBER: 22475941 Beneficial effects of a glyburide/ TITLE:

metformin combination preparation in type 2

diabetes mellitus.

AUTHOR: Bokhari Syed U; Gopal Usha M; Duckworth William C

CORPORATE SOURCE: Carl T. Hayden VA Medical Center, Phoenix, Arizona 85012,

USA.. syed.bokhari2@med.va.gov

AMERICAN JOURNAL OF THE MEDICAL SCIENCES, (2003 Feb) 325 SOURCE:

Journal code: 0370506. ISSN: 0002-9629.

PUB. COUNTRY: United States DOCUMENT TYPE: (CLINICAL TRIAL)

Journal; Article; (JOURNAL ARTICLE)

(RANDOMIZED CONTROLLED TRIAL)

LANGUAGE:

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 200303

ENTRY DATE: Entered STN: 20030221

> Last Updated on STN: 20030331 Entered Medline: 20030328

BACKGROUND: Type 2 diabetes mellitus is characterized by both insulin deficiency and insulin resistance. Effective treatment often requires therapy directed at both abnormalities. Patients on monotherapy might benefit from a combination agent such as glyburide/ metformin, which increases insulin secretion and reduces insulin resistance. METHODS: All patients taking a glyburide/ metformin preparation at the Carl T. Hayden VAMC were identified from pharmacy records. Patients with documented hemoglobin A values within 31 weeks prior and between 3 and 33 weeks after initiation of therapy (92 subjects) were examined. RESULTS: Glyburide/ metformin combination therapy reduced hemoglobin A levels from 0.087 to 0.083 (P < 0.06). Significant reductions were seen in those patients with initial levels higher than 0.08 (0.094 to 0.087; P < 0.01). No significant reductions were seen in those patients with initial levels. lower than 0.08. CONCLUSIONS: In patients on monotherapy or on dual oral therapy with inadequate control, changing to a glyburide/ metformin combination preparation may improve glucose control.

ACCESSION NUMBER: 2001408485 MEDLINE

DOCUMENT NUMBER: 21082683 PubMed ID: 11460818

TITLE: Oral antidiabetic treatment in patients with coronary

disease: time-related increased mortality on combined

glyburide/metformin therapy over a

7.7-year follow-up.

AUTHOR: Fisman E Z; Tenenbaum A; Boyko V; Benderly M; Adler Y;

Friedensohn A; Kohanovski M; Rotzak R; Schneider H; Behar

S; Motro M

CORPORATE SOURCE: Cardiac Rehabilitation Institute, the Chaim Sheba Medical

Center, Tel-Hashomer, Israel.

SOURCE: CLINICAL CARDIOLOGY, (2001 Feb) 24 (2) 151-8.

Journal code: 7903272. ISSN: 0160-9289.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200107

ENTRY DATE: Entered STN: 20010723

Last Updated on STN: 20021018 Entered Medline: 20010719

BACKGROUND: A sulfonylurea -- usually glyburide -- plus metformin constitute AB the most widely used oral antihyperglycemic combination in clinical practice. Both medications present undesirable cardiovascular effects. The issue whether the adverse effects of each of these pharmacologic agents may be additive and detrimental to the prognosis for coronary patients has not yet been specifically addressed. HYPOTHESIS: This study was designed to examine the survival in type 2 diabetics with proven coronary artery disease (CAD) receiving a combined glyburide/ metformin antihyperglycemic treatment over a long-term follow-up period. METHODS: The study sample comprised 2,275 diabetic patients, aged 45-74 years, with proven CAD, who were screened but not included in the bezafibrate infarction prevention study. In addition, 9,047 nondiabetic patients with CAD represented a reference group. Diabetics were divided into four groups on the basis of their therapeutic regimen: diet alone (n = 990), glyburide (n = 953), metformin (n = 79), and a combination of the latter two (n = 253). RESULTS: The diabetic groups presented similar clinical characteristics upon recruitment. Crude mortality rate after a 7.7-year follow-up was lower in nondiabetics (14 vs. 31.6%, p<0.001). Among diabetics, 720 patients died: 260 on diet (mortality 26.3%), 324 on glyburide (34%), 25 on metformin alone (31.6%), and 111 patients (43.9%) on combined treatment (p<0.000001). Time-related mortality was almost equal for patients on metformin and on combined therapy over an intermediate follow-up period of 4 years (survival rates 0.80 and 0.79, respectively). The group on combined treatment presented the worst prognosis over the long-term follow-up, with a time-related survival rate of 0.59 after 7 years, versus 0.68 and 0.70 for glyburide and metformin, respectively. After adjustment to variables for prognosis, the use of the combined treatment was associated with an increased hazard ratio (HR) for all-cause mortality of 1.53 (95% confidence interval [CI] 1.20-1.96), whereas glyburide and metformin alone yielded HR 1.22 (95% CI 1.02-1.45) and HR 1.26 (95% CI 0.81-1.96), respectively. Conclusions: We conclude that after a 7.7-year follow-up, monotherapy with either glyburide or metformin in diabetic patients with CAD yielded a similar outcome and was associated with a modest increase in mortality. However, time-related mortality was markedly increased when a combined glyburide/ metformin treatment was used.

L11 ANSWER 23 OF 56 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: DOCUMENT NUMBER:

2001:448868 BIOSIS

PREV200100448868

TITLE:

Durable antidiabetic effect of glyburide/

metformin tablets as initial therapy for type 2

diabetes.

AUTHOR(S):

Garber, Alan J. (1); Piper, Beth Ann (1); Park, Jong-Soon

(1)

CORPORATE SOURCE:

(1) Houston, TX USA

SOURCE:

Diabetes, (June, 2001) Vol. 50, No. Supplement 2, pp. A113.

Meeting Info.: 61st Scientific Sessions of the American Diabetes Association Philadelphia, Pennsylvania, USA June

22-26, 2001 ISSN: 0012-1797.

DOCUMENT TYPE:

Conference English

LANGUAGE:

SUMMARY LANGUAGE: English

L11 ANSWER 24 OF 56 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: DOCUMENT NUMBER:

2001:448854 BIOSIS PREV200100448854

TITLE:

Durable antidiabetic effect of glyburide/

metformin tablets as second-line therapy for type 2

diabetes.

AUTHOR(S):

Blonde, Lawrence (1); Rosenstock, Julio (1); Piper, Beth

Ann (1); Henry, David (1)

CORPORATE SOURCE:

(1) New Orleans, LA USA

SOURCE:

Diabetes, (June, 2001) Vol. 50, No. Supplement 2, pp. A106.

print.

Meeting Info.: 61st Scientific Sessions of the American Diabetes Association Philadelphia, Pennsylvania, USA June

22-26, 2001

ISSN: 0012-1797.

DOCUMENT TYPE:

Conference English

LANGUAGE: SUMMARY LANGUAGE:

English

ACCESSION NUMBER: 2001061324 MEDLINE

DOCUMENT NUMBER: 20530757 PubMed ID: 11077467

TITLE: Glyburide/metformin (Glucovance) for

type 2 diabetes.

AUTHOR: Anonymous

SOURCE: MEDICAL LETTER ON DRUGS AND THERAPEUTICS, (2000 Nov 13) 42

(1092) 105-6.

Journal code: 2985240R. ISSN: 0025-732X.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH: 200012

ENTRY DATE: Entered STN: 20010322

Last Updated on STN: 20021018 Entered Medline: 20001228

L11 ANSWER 31 OF 56 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2002:580061 BIOSIS DOCUMENT NUMBER: PREV200200580061

TITLE: Efficacy of glyburide/metformin tablets

versus metformin plus rosiglitazone in patients with type 2

diabetes inadequately controlled with metformin

monotherapy.

AUTHOR(S): Mohideen, P. (1); Klein, E.; Bruce, S. (1)

CORPORATE SOURCE: (1) Bristol-Myers Squibb Pharmaceutical Research Institute,

Princeton, NJ USA

SOURCE: Diabetologia, (August, 2002) Vol. 45, No. Supplement 2, pp.

A 242. print.

Meeting Info.: 38th Annual Meeting of the European Association for the Study of Diabetes (EASD) Budapest, Hungary September 01-05, 2002 European Association for the

Study of Diabetes . ISSN: 0012-186X.

DOCUMENT TYPE: Conference LANGUAGE: English

L11 ANSWER 32 OF 56 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2001:449039 BIOSIS DOCUMENT NUMBER: PREV200100449039

TITLE: Combination therapy in type 2 diabetes:

Repaglinide/metformin vs glyburide/

metformin.

AUTHOR(S): Jinagouda, Sujata (1); Schwartz, Sherwyn; Huffman, David;

Weinstein, Richard; Davidson, Jaime; Huang, Wonchin;

Reinhardt, Rickey

CORPORATE SOURCE: (1) Alhambra, CA USA

SOURCE: Diabetes, (June, 2001) Vol. 50, No. Supplement 2, pp. A439.

print.

Meeting Info.: 61st Scientific Sessions of the American Diabetes Association Philadelphia, Pennsylvania, USA June

22-26, 2001 ISSN: 0012-1797.

DOCUMENT TYPE: Conference LANGUAGE: English SUMMARY LANGUAGE: English

L11 ANSWER 33 OF 56 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2001:449020 BIOSIS DOCUMENT NUMBER: PREV200100449020

TITLE: 20-month durability of glyburide/

metformin tablets on glycemic control as initial

therapy for type 2 diabetes.

AUTHOR(S): Donovan, Daniel (1); Piper, Beth Ann (1); Park, Jong-Soon

CORPORATE SOURCE:

(1) New York, NY USA

SOURCE:

Diabetes, (June, 2001) Vol. 50, No. Supplement 2, pp. A434.

print.

Meeting Info.: 61st Scientific Sessions of the American Diabetes Association Philadelphia, Pennsylvania, USA June

22-26, 2001 ISSN: 0012-1797.

DOCUMENT TYPE: LANGUAGE:

Conference English

SUMMARY LANGUAGE:

English

L11 ANSWER 34 OF 56 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V. on STN

ACCESSION NUMBER:

2002269883 EMBASE

TITLE:

Erratum: 'Simultaneous glyburide/

metformin therapy is superior to component

monotherapy as an initial pharmacological treatment for type 2 diabetes' (Diabetes, Obesity and Metabolism vol. 4

(3) (201-208)).

SOURCE:

Diabetes, Obesity and Metabolism, (2002) 4/4 (286).

ISSN: 1462-8902 CODEN: DOMEF6

COUNTRY:

United Kingdom Journal; Errata

DOCUMENT TYPE: FILE SEGMENT:

Endocrinology 003

LANGUAGE:

English

L11 ANSWER 35 OF 56 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V. on STN

ACCESSION NUMBER:

2000404460 EMBASE

TITLE:

Transitioning patients with type 2 diabetes to a fixed

combination > glyburide/metformin

tablet.

AUTHOR:

Blonde L.; Sandberg M.I.

CORPORATE SOURCE:

Dr. L. Blonde, Department of Internal Medicine, Ochsner Diabetes Clinical, Research Unit, 1514 Jefferson Highway, New Orleans, LA 70121, United States. lblonde@Ochsner.org Diabetes Technology and Therapeutics, (2000) 2/3 (479-480).

SOURCE:

Refs: 3

ISSN: 1520-9156 CODEN: DTTHFH

COUNTRY:

United States

DOCUMENT TYPE:

Journal; Letter

FILE SEGMENT:

Endocrinology 003 037 Drug Literature Index

LANGUAGE:

English

L11 ANSWER 43 OF 56 MEDLINE ON STN ACCESSION NUMBER: 2003397110 MEDLINE

DOCUMENT NUMBER: 22815632 PubMed ID: 12934950

TITLE: Using the electronic medical record to enhance the use of

combination drugs.

AUTHOR: Wells Brian J; Lobel Keith D; Dickerson Lori M

CORPORATE SOURCE: Department of Family Medicine, Medical University of South

Carolina, Charleston, SC 29425, USA.. wellsbj@musc.edu

SOURCE: AMERICAN JOURNAL OF MEDICAL QUALITY, (2003 Jul-Aug) 18 (4)

147-9.

Journal code: 9300756. ISSN: 1062-8606.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200309

ENTRY DATE: Entered STN: 20030826

Last Updated on STN: 20030910 Entered Medline: 20030909

The objective of this study was to increase combination drug prescriptions AB through the use of electronic point-of-care reminders, thereby maintaining quality while decreasing medication costs. The electronic medical record (EMR) was used to identify all patients who were potential candidates for one of the following 3 currently available combination drugs: fluticasone-salmeterol, amlodipine-benazepril, or glyburidemetformin. Point-of-care electronic reminders were attached to the medication record of the EMR for each patient, and providers were asked to consider using the available combination medication. Of the patients who had electronic reminders attached to their charts and were seen at the clinic during the study period, 47 of 175 were switched to a combination medication. A cost-savings analysis showed a total annual savings of dollars 6,159.30. Point-of-care reminders are a simple and effective tool for quality-improvement interventions. Combination drugs may play an important role in controlling medication costs.

ACCESSION NUMBER: 2001376630 EMBASE

Trends in diabetes care. TITLE:

AUTHOR: Haveles E.B.

CORPORATE SOURCE: Prof. E.B. Haveles, Old Dominion University, Norfolk, Va,

United States

Drug Topics, (1 Oct 2001) 145/19 SUPPL. (29s-36s). SOURCE:

ISSN: 0012-6616 CODEN: DGTNA7

COUNTRY: United States

Journal; General Review DOCUMENT TYPE: Endocrinology FILE SEGMENT: 003 Internal Medicine 006

Public Health, Social Medicine and Epidemiology 017

037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

Type 2 diabetes is a common cause of morbidity and mortality that can be prevented or delayed with glycemic control. A sequential approach to treating Type 2 diabetes - initiating monotherapy and moving to combination therapy when monotherapy fails - is widely used and accepted. Sulfonylureas can undoubtedly improve glycemic control with initial therapy and later with the addition of other antidiabetic medications. Metformin is also an option as either monotherapy or combination therapy. Based upon the results of the UKPDS, metformin may be of benefit for significantly obese patients because of the lack of weight gain. In fact, patients may actually lose weight while on metformin. Acarbose may be an option for patients with elevated lipid levels. Acarbose may actually improve the lipid profile by reducing the ratio of LDL-to-HDL cholesterol. The thiazolidinediones have not been shown to have a consistent effect on lipid levels, and these agents cause weight gain. No studies are available that evaluate the effects of repaglinide on lipid levels. There is debate regarding initiating monotherapy or combination therapy as the first-line approach to treating Type 2 diabetes. The ADA continues to recommend sulfonylureas as appropriate monotherapy for initially treating Type 2 diabetes. Eventually, most patients will require some form of combination antidiabetic therapy. Most research involves metformin complemented by a sulfonylurea, though other antidiabetic combinations have been used with success. Glyburide/metformin fixed combination is now available, which may improve patient compliance because the patient must remember to take only one "drug," not two separate drugs. However, patients are locked into specific doses, which can create problems. Use of two separate medications in combination affords the clinician the ability to change the dose of one medication at a time and observe for results. Glipizide-GITS, whether as monotherapy or in combination with metformin, is a new option in treating Type 2 diabetes. The formulation is well tolerated, appears to mimic natural insulin release, and is a true once-daily dose form as either first-line or combination therapy. It provides 24-hour control, which is not only convenient but also improves patient compliance. Glipizide-GITS lowers fasting insulin levels more than glyburide and immediate-release glipizide, and long-term data show no weight gain on average and also the possibility that it may actually lower plasma lipid and triglyceride levels. If combination therapy is necessary, the addition of another antidiabetic drug to glipizide-GITS continues to lower HbA(1c) levels. Lastly, efforts to improve patient compliance, continuous monitoring of plasma glucose levels and HbA(1c) levels, and optimizing antidiabetic therapy can improve patient outcomes.

ACCESSION NUMBER: 2002216237 MEDLINE

DOCUMENT NUMBER: 21948517 PubMed ID: 11952029

TITLE: Adherence to oral antidiabetic therapy in a managed care

organization: a comparison of monotherapy, combination

therapy, and fixed-dose combination therapy.

AUTHOR: Melikian Caron; White T Jeffrey; Vanderplas Ann; Dezii

Christopher M; Chang Eunice

CORPORATE SOURCE: Prescription Solutions, Costa Mesa, California 92626, USA..

caron.melikian@phs.com

SOURCE: CLINICAL THERAPEUTICS, (2002 Mar) 24 (3) 460-7.

Journal code: 7706726. ISSN: 0149-2918.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200210

ENTRY DATE: Entered STN: 20020416

Last Updated on STN: 20021002 Entered Medline: 20021001

BACKGROUND: Although medication adherence is one of the most important AB aspects of the management of diabetes mellitus, low rates of adherence have been documented. OBJECTIVE: This study sought to examine medication adherence among patients with diabetes mellitus in a managed care organization who were receiving antidiabetic monotherapy (metformin or glyburide), combination therapy (metformin and glyburide), or fixed-dose combination therapy (glyburide/metformin). METHODS: Medication adherence was evaluated through a retrospective database analysis of pharmacy claims. The adherence rate was defined as the sum of the days' supply of oral antidiabetic medication obtained by the patient during the follow-up period divided by the total number of days in the designated follow-up period (180 days). Health plan members were included in the analysis if they had an index pharmacy claim for an oral antidiabetic medication between August 1 and December 31, 2000, were continuously enrolled in the health plan, and were aged > or =18 years. A 6-month pre-index period was used to classify patients as newly treated or previously treated. Patients were grouped according to their medication-use patterns. RESULTS: After adjustment for potential confounding factors, including overall medication burden at index, there were no significant differences in adherence rates among 6502 newly treated patients receiving monotherapy, combination therapy, or fixed-dose combination therapy. Among the 1815 previously treated patients receiving glyburide or metformin monotherapy who required the addition of the alternative agent, resulting in combination therapy, adherence rates were significantly lower (54.0%; 95% CI, 0.52-0.55) than in the 105 patients receiving monotherapy who were switched to fixed-dose combination therapy (77.0%; 95% CI, 0.72-0.82). The 59 previously treated patients receiving combination therapy who were switched to fixed-dose combination therapy had a significant improvement in adherence after the switch (71.0% vs 87.0%; P < 0.001). CONCLUSIONS: In a managed care organization, previously treated patients receiving monotherapy with an oral antidiabetic medication who required additional therapy exhibited significantly greater adherence when they were switched to fixed-dose combination therapy compared with combination therapy. Patients receiving combination therapy who were switched to fixed-dose combination therapy exhibited significantly greater adherence after the switch.

ACCESSION NUMBER: 2003397110 MEDLINE

DOCUMENT NUMBER: 22815632 PubMed ID: 12934950

TITLE: Using the electronic medical record to enhance the use of

combination drugs.

AUTHOR: Wells Brian J; Lobel Keith D; Dickerson Lori M

CORPORATE SOURCE: Department of Family Medicine, Medical University of South

Carolina, Charleston, SC 29425, USA.. wellsbj@musc.edu

SOURCE: AMERICAN JOURNAL OF MEDICAL QUALITY, (2003 Jul-Aug) 18 (4)

147-9.

Journal code: 9300756. ISSN: 1062-8606.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200309

ENTRY DATE: Entered STN: 20030826

Last Updated on STN: 20030910 Entered Medline: 20030909

The objective of this study was to increase combination drug prescriptions through the use of electronic point-of-care reminders, thereby maintaining quality while decreasing medication costs. The electronic medical record (EMR) was used to identify all patients who were potential candidates for one of the following 3 currently available combination drugs: fluticasone-salmeterol, amlodipine-benazepril, or glyburide-metformin. Point-of-care electronic reminders were attached to the medication record of the EMR for each patient, and providers were asked to consider using the available combination medication. Of the patients who had electronic reminders attached to their charts and were seen at the clinic during the study period, 47 of 175 were switched to a combination medication. A cost-savings analysis showed a total annual savings of dollars 6,159.30. Point-of-care reminders are a simple and effective tool for quality-improvement interventions. Combination drugs may play an important role in controlling medication costs.

```
ANSWER 1 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN
L1
     338752-31-1 REGISTRY
RN
     Benzamide, 5-chloro-N-[2-[4-[[[(cyclohexylamino)carbonyl]amino]sulfonyl]ph
CN
     enyl]ethyl]-2-methoxy-, mixt. with N,N-dimethylimidodicarbonimidic diamide
     monohydrochloride (9CI)
                               (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Imidodicarbonimidic diamide, N,N-dimethyl-, monohydrochloride, mixt.
CN
     contg. (9CI)
OTHER NAMES:
     Glucovance
CN
CN
     Glyburide-metformin hydrochloride mixt.
     C23 H28 Cl N3 O5 S . C4 H11 N5 . Cl H
MF
CI
     MXS
SR
     CA
                  BIOSIS, CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL
LC
     STN Files:
     CM
          1
     CRN
         10238-21-8
     CMF C23 H28 C1 N3 O5 S
  OMe
         ~ NH— СН<sub>2</sub>— СН<sub>2</sub>
                                   NH-C-
                                         – NH
     CM
          2
         1115-70-4 (657-24-9)
     CRN
     CMF C4 H11 N5 . Cl H
```

● HCl

- 8 REFERENCES IN FILE CA (1937 TO DATE)
- 8 REFERENCES IN FILE CAPLUS (1937 TO DATE)
- L1 ANSWER 2 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN
- RN 338752-30-0 REGISTRY
- CN Benzamide, 5-chloro-N-[2-[4-[[[(cyclohexylamino)carbonyl]amino]sulfonyl]ph enyl]ethyl]-2-methoxy-, mixt. with N,N-dimethylimidodicarbonimidic diamide (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Imidodicarbonimidic diamide, N,N-dimethyl-, mixt. contg. (9CI) OTHER NAMES:

CN Glyburide-metformin mixt.

MF $\tilde{}$ C23 H28 Cl N3 O5 S . C4 H11 N5

CI MXS

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL

CM 1

CRN 10238-21-8

CMF C23 H28 C1 N3 O5 S

CM 2

CRN 657-24-9 CMF C4 H11 N5

4 REFERENCES IN FILE CA (1937 TO DATE)

4 REFERENCES IN FILE CAPLUS (1937 TO DATE)

L1 ANSWER 3 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN

RN 88159-36-8 REGISTRY

CN Sodium glipizide

MF C21 H27 N5 O4 S . Na

LC STN Files: CA, CAPLUS, DRUGPAT

CRN (29094-61-9)

■ N=

1 REFERENCES IN FILE CA (1937 TO DATE)

1 REFERENCES IN FILE CAPLUS (1937 TO DATE)

L1 ANSWER 4 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN

RN 58840-24-7 REGISTRY

CN 4-Pyrimidinecarboxylic acid, 1,2,3,6-tetrahydro-2,6-dioxo-, compd. with

N,N-dimethylimidodicarbonimidic diamide (1:1) (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES:

CN Imidodicarbonimidic diamide, N,N-dimethyl-, mono(1,2,3,6-tetrahydro-2,6-dioxo-4-pyrimidinecarboxylate) (9CI)

OTHER NAMES:

CN

Metformin orotate

MF C5 H4 N2 O4 . C4 H11 N5

LC STN Files: BEILSTEIN*, CA, CAPLUS, RTECS*

(*File contains numerically searchable property data)

CM 1

CRN 657-24-9 CMF C4 H11 N5

· CM 2

CRN 65-86-1 CMF C5 H4 N2 O4

$$\begin{array}{c|c} O & H & CO_2H \\ \hline HN & . \\ \hline \\ O & \end{array}$$

1 REFERENCES IN FILE CA (1937 TO DATE).

1 REFERENCES IN FILE CAPLUS (1937 TO DATE)

L1 ANSWER 5 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN

RN 53950-18-8 REGISTRY

CN Propanoic acid, 2-(4-chlorophenoxy)-2-methyl-, compd. with N,N-dimethylimidodicarbonimidic diamide (1:1) (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES:

CN Imidodicarbonimidic diamide, N,N-dimethyl-, mono[2-(4-chlorophenoxy)-2-methylpropanoate] (9CI)

OTHER NAMES:

CN ANP 4324

CN Metformin clofibrate

MF C10 H11 Cl O3 . C4 H11 N5

LC STN Files: BEILSTEIN*, BIOSIS, CA, CAPLUS, IFICDB, IFIPAT, IFIUDB (*File contains numerically searchable property data)

CM 1

CRN 882-09-7

CMF C10 H11 Cl O3

CM 2

CRN 657-24-9 CMF C4 H11 N5

4 REFERENCES IN FILE CA (1937 TO DATE)

4 REFERENCES IN FILE CAPLUS (1937 TO DATE)

L1 ANSWER 6 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN

RN 51394-30-0 REGISTRY

CN Benzenesulfonamide, 4-chloro-N-[(propylamino)carbonyl]-, mixt. with N,N-dimethylimidodicarbonimidic diamide (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES:

CN Imidodicarbonimidic diamide, N,N-dimethyl-, mixt. contg. (9CI) OTHER NAMES:

CN Chlorpropamide-1,1-dimethylbiguanide mixt.

CN Chlorpropamide-metformin mixt.

CN Obinese

MF C10 H13 Cl N2 O3 S . C4 H11 N5

CI MXS

LC STN Files: BEILSTEIN*, CA, CAPLUS, EMBASE (*File contains numerically searchable property data)

CM 1

CRN 657-24-9 CMF C4 H11 N5

CM 2

CRN 94-20-2

CMF C10 H13 Cl N2 O3 S

2 REFERENCES IN FILE CA (1937 TO DATE)
2 REFERENCES IN FILE CAPLUS (1937 TO DATE)

L1 ANSWER 7 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN

RN 38950-16-2 REGISTRY

CN Benzenesulfonamide, N-[(butylamino)carbonyl]-4-methyl-, compd. with N,N-dimethylimidodicarbonimidic diamide (1:1) (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES:

CN Biguanide, 1,1-dimethyl-, compd. with 1-butyl-3-p-tolylsulfonylurea (6CI)

CN Imidodicarbonimidic diamide, N,N-dimethyl-, compd. with

N-[(butylamino)carbonyl]-4-methylbenzenesulfonamide (1:1) (9CI)

CN Urea, 1-butyl-3-(p-tolylsulfonyl)-, compd. with 1,1-dimethylbiguanide (7CI)

OTHER NAMES:

CN 1-Butyl-3-(p-tolylsulfonyl)urea and 1,1-dimethylbiguanide adduct

CN Metformin tolbutamide salt

MF C12 H18 N2 O3 S . C4 H11 N5

LC STN Files: BEILSTEIN*, CA, CAOLD, CAPLUS, TOXCENTER (*File contains numerically searchable property data)

CM 1

CRN 657-24-9 CMF C4 H11 N5

CM 2

CRN 64-77-7

CMF C12 H18 N2 O3 S

- 4 REFERENCES IN FILE CA (1937 TO DATE)
- 4 REFERENCES IN FILE CAPLUS (1937 TO DATE)
- 2 REFERENCES IN FILE CAOLD (PRIOR TO 1967)
- L1 ANSWER 8 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN
- RN 34461-22-8 REGISTRY
- CN 2-Naphthalenecarboxylic acid, 4,4'.-methylenebis[3-hydroxy-, compd. with N,N-dimethylimidodicarbonimidic diamide (1:2) (9CI) (CA INDEX NAME)

```
OTHER CA INDEX NAMES:
     2-Naphthoic acid, 4,4'-methylenebis[3-hydroxy-, compd. with
     1,1-dimethylbiguanide (1:2) (8CI)
     Imidodicarbonimidic diamide, N,N-dimethyl-, 4,4'-methylenebis[3-hydroxy-2-
CN
     naphthalenecarboxylate] (2:1) (9CI)
OTHER NAMES:
CN
     Metformin pamoate
MF
     C23 H16 O6 . 2 C4 H11 N5
                  BIOTECHNO, CA, CAPLUS, CHEMLIST, CSCHEM, EMBASE, MRCK*,
LC
     STN Files:
       USPATFULL
         (*File contains numerically searchable property data)
                      EINECS**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
     CM
     CRN
          657-24-9
     CMF
          C4 H11 N5
     NH
            NH
Me_2N-C-NH-C-NH_2
     CM
     CRN
          130-85-8
     CMF
         C23 H16 O6
              CO2H
              OH
         CH<sub>2</sub>
  HO
HO2C
               5 REFERENCES IN FILE CA (1937 TO DATE)
               5 REFERENCES IN FILE CAPLUS (1937 TO DATE)
L1
     ANSWER 9 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN
RN
     29094-61-9 REGISTRY
     Pyrazinecarboxamide, N-[2-[4-[[[(cyclohexylamino)carbonyl]amino]sulfonyl]p
CN
     henyl]ethyl]-5-methyl- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Urea, 1-cyclohexyl-3-[[p-[2-(5-methylpyrazinecarboxamido)ethyl]phenyl]sulf
     onyl] - (8CI)
OTHER NAMES:
CN
     Aldiab
CN
     CP 28720
CN
     Digrin
CN
     Dipazide
CN
     Glibenese
CN
     Glibetin
CN
     Glican
```

Glidiab

CN

```
Glipid
CN
     Glipizide
CN
CN
     Gluco-Rite
     Glucolip
CN
     Glucotrol
CN
CN
     Glucotrol Xl
     Glucozide
CN
     Glupitel
CN
CN
     Glupizide
CN
     Glvde
     Glydiazinamide
CN
CN
     Glynase
     K 4024
CN
CN
     Melizide
CN
     Mindiab
CN
     Minidab
CN
     Minidiab
CN
     Minodiab
     N-(4-[.beta.-(5-Methylpyrazine-2-carboxamido)ethyl]benzenesulfonyl)-N'-
CN
     cyclohexylurea
CN
     Napizide
     Ozidia
CN
CN
     Sucrazide
CN
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FS
     3D CONCORD
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       IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, PHAR, PHARMASEARCH, PROMT, RTECS*,
       SPECINFO, TOXCENTER, USAN, USPAT2, USPATFULL, VETU
         (*File contains numerically searchable property data)
     Other Sources:
                      EINECS**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
                                           - C- NH
                                       NH-
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PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

592 REFERENCES IN FILE CA (1937 TO DATE)
3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
594 REFERENCES IN FILE CAPLUS (1937 TO DATE)

L1 ANSWER 10 OF 11 REGISTRY COPYRIGHT 2003 ACS on STN
RN 1115-70-4 REGISTRY
CN Imidodicarbonimidic diamide, N,N-dimethyl-, monohydrochloride (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
CN Biguanide, 1,1-dimethyl-, hydrochloride (6CI)
CN Biguanide, 1,1-dimethyl-, monohydrochloride (8CI)
OTHER NAMES:

CN 1,1-Dimethylbiguanide hydrochloride

CN Apophage CN Benofomin CN Dabex

```
CN
     Denkaform
CN
     Dextin
     Diabefagos
CN
CN
     Diabetmin
CN:
     Diabetosan
CN
     Diabex
     Diaformin
CN
CN
     Dialon
CN
     Diformin
CN
     Diformin Retard
CN
     Dimefor
CN
     Fornidd
CN
     Geamet
CN
     Glucaminol
CN
     Glucofago
CN
     Glucoform
CN
     Glucomet
CN
     Glucomin
CN
     Glucomine
CN
     Gluconil
CN
     Glucophage
CN
     Glucophage 850
CN
     Glucophage Forte
CN
     Glucophage Retard
CN
     Glucophage-Mite
CN
     Gludepatic
CN
     Glufor
CN
     Gluformin
     Glumeformin
CN
CN
     Glumin
CN
     Glupermin
CN
     Glyceriphage
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    Glyciphage
CN
     Glycon
CN
     Glyformin
CN
     LA 6023
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     Mequan
CN
     Metforal
CN
     Metformin hydrochloride
CN
     Metomin
CN
     Miformin
CN
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CN
     N1,N1-Dimethylbiguanide hydrochloride
CN
     Orabet
ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
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     C4 H11 N5 . Cl H
CI
     COM
                  ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*,
LC
     STN Files:
       BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CABA, CAOLD, CAPLUS, CASREACT, CBNB,
       CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, DIOGENES, DRUGUPDATES,
       EMBASE, GMELIN*, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MRCK*, MSDS-OHS,
       PHAR, PHARMASEARCH, PROMT, RTECS*, TOXCENTER, ULIDAT, USAN, USPAT2,
       USPATFULL
         (*File contains numerically searchable property data)
                      EINECS**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
CRN
     (657-24-9)
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19 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

ANSWER 150 OF 183 USPATFULL on STN

ACCESSION NUMBER:

2003:24185 USPATFULL

TITLE:

Combination therapy for type II diabetes or Syndrome X

INVENTOR(S):

Gwynne, John Thomas, Doylestown, PA, UNITED STATES

Vitou, Philippe John Robert, Paris, FRANCE

Randazzo, Bruce Paul, Rydal, PA, UNITED STATES

PATENT ASSIGNEE(S):

Wyeth, Madison, NJ (U.S. corporation)

NUMBER KIND DATE -----

PATENT INFORMATION:

US 2003018028 A1 20030123

APPLICATION INFO.:

US 2002-163707 Al 20020606 (10)

> NUMBER DATE

PRIORITY INFORMATION:

US 2001-296502P

20010607 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Arnold S. Milowsky, 5 Giralda Farms, Madison, NJ, 07940

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: LINE COUNT:

1108

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention provides methods of using a pharmacological combination of a biguanide agents, such as metformin, and one or more PTPase inhibiting agents and, optionally, one or more sulfonlylurea agents, including glyburide, glyburide, glipizide, glimepiride, chlorpropamide, tolbutamide, or tolazamide, for treatment in a mammal of Syndrome X, type II diabetes or metabolic disorders mediated by insulin resistance or hyperglycemia. Further included in this invention is a method of modulating blood glucose levels in a mammal utilizing the combination of one or more PTPase inhibiting agents and one or more sulfonlylurea agents.

ANSWER 162 OF 183 USPATFULL on STN

ACCESSION NUMBER:

1999:81839 USPATFULL

TITLE:

Methods for use of cryptolepine analogs with

hypoglycemic activity

INVENTOR (S):

Bierer, Donald E., Daly City, CA, United States

PATENT ASSIGNEE(S):

Shaman Pharmaceuticals, Inc., South San Francisco, CA,

United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION:

_______ US 5925647 19990720

APPLICATION INFO.:

19971020 (8) US 1997-955320

RELATED APPLN. INFO.:

Division of Ser. No. US 1995-484424, filed on 7 Jun

1995, now patented, Pat. No. US 5681958

DOCUMENT TYPE:

Utility

FILE SEGMENT:

Granted

PRIMARY EXAMINER: LEGAL REPRESENTATIVE:

Schenkman, Leonard Pennie & Edmonds LLP

NUMBER OF CLAIMS:

1,4

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

9 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT:

3932

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Novel cryptolepine analogs useful as hypoglycemic agents and methods for their use as hypoglycemic agents, for example, in the treatment of diabetes, and a method for their synthesis are described. As hypoglycemic agents, the novel cryptolepine analogs are useful for the treatment of insulin-dependent diabetes mellitus (IDDM or Type I) and non-insulin dependent diabetes mellitus (NIDDM or Type II).

L19 ANSWER 1 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:315117 USPATFULL

TITLE: ANTIDIABETIC FORMULATION AND METHOD

INVENTOR(S): PIPER, BETH ANNE, HOPEWELL, NJ, UNITED STATES

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: STEPHEN B. DAVIS, BRISTOL-MYERS SQUIBB COMPANY, PATENT

DEPARTMENT, P O BOX 4000, PRINCETON, NJ, 08543-4000

NUMBER OF CLAIMS: 76 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 10 Drawing Page(s)

LINE COUNT: 1927

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A low dose antidiabetic pharmaceutical formulation is provided, especially adapted for treating Type II diabetes in drug naive patients, which includes a combination of metformin (employed in a reduced amount (less than 800 mg metformin per day) compared to that employed in generally accepted medical practice) and at least one other antidiabetic agent such as a sulfonyl urea, for example, qlyburide, which combination provides at least about substantially equivalent efficacy in treating diabetes in drug naive patients, as do antidiabetic formulations containing metformin employed in dosages prescribed in generally accepted medical practice for first line therapy in treating diabetes, but with substantially reduced side effects, such as hypoglycemia and/or gastrointestinal distress. A method for treating diabetes in drug naive human patients is also provided employing the above formulation to reduce insulin resistance and/or post-prandial glucose excursion and/or hemoglobin 1Ac, and/or increase post-prandial insulin, thereby treating the diabetes.

L19 ANSWER 2 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2000:24678 USPATFULL

TITLE: Salts of metformin and method

Timmins, Peter, Merseyside, United Kingdom INVENTOR (S): Winter, William J., Lebanon, NJ, United States

Srivastava, Sushil K., Dayton, NJ, United States Bretnall, Alison E., Chester, United Kingdom

Wei, Chenkou, Princeton Junction, NJ, United States Powers, Gerald L., North Brunswick, NJ, United States Bristol-Myers Squibb Company, Princeton, NJ, United

States (U.S. corporation)

NUMBER KIND DATE ______

PATENT INFORMATION: US 6031004 20000229

US 1999-262526 19990304 (9) APPLICATION INFO.:

Continuation of Ser. No. US 1997-986586, filed on 8 Dec RELATED APPLN. INFO.:

DOCUMENT TYPE: Utility FILE SEGMENT: Granted Fay, Zohreh PRIMARY EXAMINER: LEGAL REPRESENTATIVE: Rodney, Burton

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT: 651

PATENT ASSIGNEE(S):

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Novel salts of the antidiabetic agent metformin acre provided which are metformin salts of dibasic acids (2:1 molar ratio), preferably metformin (2:1) fumarate and metformin

(2:1) succinate, which may be employed alone or in combination with another antihyperglycemic agent such as glyburide, for treating diabetes. A method for treating diabetes employing the novel metformin salt by itself or in combination with another

antidiabetic agent is also provided.

2003:112605 USPATFULL ACCESSION NUMBER:

Formulations for the prevention and treatment of TITLE:

insulin resistance and type 2 diabetes mellitus

Richardson, Kenneth T., Anchorage, AK, UNITED STATES INVENTOR (S):

Pearson, Don C., Lakewood, WA, UNITED STATES

PATENT ASSIGNEE(S):

ChronoRX LLC, Anchorage, AK (U.S. corporation)

NUMBER KIND DATE ----- ----- ----- -----

PATENT INFORMATION: US 2003077335 A1 20030424

APPLICATION INFO.: US 2001-33730 A1 20011102 (10)

> NUMBER DATE -----

PRIORITY INFORMATION: US 2000-245471P 20001103 (60)

> US 2000-245950P 20001103 (60)

US 2000-256033P 20001213 (60)

DOCUMENT TYPE: Utility

APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO

CENTER, EIGHTH FLOOR, SAN FRANCISCO, CA, 94111-3834

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1

4450 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The compositions and dosage forms of the invention are clinically useful as methods for increasing the effectiveness, efficiency and safety of biguanides (metformin) and/or sulfonylureas in the prevention and treatment of insulin resistance and diabetes mellitus, alone or in combination, as a nutrient for humans. The carefully chosen active ingredients of the invention are designed in a modular fashion to prevent and rectify adverse events associated with insulin resistance syndrome and diabetes mellitus, and with the clinical use of biguanides (metformin) and/or the sulfonylureas. These modules are: (1) Mitochondrial Metabolic Group, (2) Plasma and Mitochondrial Membrane Integrity Group, (3) Nocturnal Group and, (4) Insulin Alternative Group. When used in concert with a biquanide, a sulfonylurea or with a combination of both, the invention will broaden the clinical usefulness of these drugs. The invention will retard the progression of insulin resistance to type 2 diabetes, and reduce the serious microvascular and macrovascular complications commonly associated with insulin resistance syndrome and diabetes mellitus.

L19 ANSWER 6 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1988:156419 HCAPLUS

DOCUMENT NUMBER: 108:156419

Preparation and evaluation of metformin TITLE:

hydrochloride controlled-release tablets Abdallah, O. Y.; Boraie, N. A.; Naggar, V. F.

AUTHOR (S): Fac. Pharm., Univ. Alexandria, Alexandria, Egypt CORPORATE SOURCE:

SOURCE: S.T.P. Pharma (1988), 4(1), 15-20

CODEN: STPPEF; ISSN: 0758-6922

DOCUMENT TYPE: Journal English LANGUAGE:

Metformin-HCl tablets intended for controlled release

were prepd. using Me cellulose, Et cellulose, cellulose acetate, cellulose triacetate and Eudragit RS,

RL or S. The techniques employed were direct compression, wet granulation

or copptn. followed by compression. The release properties of the resulting tablets were evaluated in 0.1N HCl and phosphate

buffer (pH 6.8). The wet granulation technique could be applied successfully with Et cellulose, Eudragit RS and Eudragit RL. Me cellulose in a matrix prepd. by copptn. showed great promise as a

retardant for release. The effect of varying the relative proportion of this polymer was also studied. The dissoln. properties of 4 com. regular

tablets and a sustained-release tablet were also detd. The release patterns were examd. from the standpoint of a

diffusion-controlled process and that of 1st-order kinetics process.

L19 ANSWER 7 OF 256 USPATFULL on STN

2003:123367 USPATFULL ACCESSION NUMBER:

Method of treating metabolic disorders especially TITLE:

diabetes, or a disease or condition associated with

Gatlin, Marjorie Regan, Hoboken, NJ, United States INVENTOR(S):

Ball, Michele Ann, Morris Plains, NJ, United States Mannion, Richard Owen, Mount Arlington, NJ, United

Karnachi, Anees Abdulguadar, Hillsborough, NJ, United

Guitard, Christiane, Hegenheim, FRANCE Allison, Malcolm, Basel, SWITZERLAND

PATENT ASSIGNEE(S): Novartis AG, Basel, SWITZERLAND (non-U.S. corporation)

> NUMBER KIND DATE ______

US 6559188 B1 20030506 US 2000-663264 20000915 PATENT INFORMATION:

APPLICATION INFO.: 20000915 (9)

> NUMBER DATE ______

US 2000-304196P 20000407 (60) PRIORITY INFORMATION:

US 2000-240918P 20000309 (60) US 1999-242911P 19990917 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

Weddington, Kevin E. PRIMARY EXAMINER: Thallemer, John D. LEGAL REPRESENTATIVE:

11 NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT: 2176

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a combination, such as a combined preparation or pharmaceutical composition, respectively, which comprises nateglinide

(I) ##STR1##

or repaglinide and at least one other antidiabetic compound selected from the group consisting of thiazolidinedione derivatives (glitazones), sulfonyl urea derivatives and metformin for simultaneous, separate or sequential use in the prevention, delay of progression or treatment of diseases, especially metabolic disorders and in particular type 2 diabetes and diseases and conditions associated with diabetes; to a composition, respectively, which comprises nateglinide and a pharmaceutically acceptable carrier and to a process of making such composition; the use of such combination or composition for the preparation of a medicament for the prevention, delay of progression or treatment of metabolic disorders; a method of prevention, delay of progression or treatment of diseases in warm-blooded animals; the use of such combination or composition for the cosmetic treatment of a mammal in order to effect a cosmetically beneficial loss of body weight; and to a method of improving the bodily appearance of a warm-blooded animal.

L19 ANSWER 10 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2000:636165 HCAPLUS

DOCUMENT NUMBER: 133:227811

Directly compressed metformin hydrochloride TITLE:

INVENTOR(S):

Kumar, Vijai

PATENT ASSIGNEE(S):

Pharmalogix, Inc., USA

SOURCE:

U.S., 9 pp.

DOCUMENT TYPE:

CODEN: USXXAM

Patent

LANGUAGE:

English

1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

KIND DATE PATENT NO. APPLICATION NO. DATE US 1998-139361 19980825 ----- ---- ---- -----US 6117451 20000912 PRIORITY APPLN. INFO.: US 1998-139361 19980825

Metformin Hydrochloride (herein referred to as metformin HCl) that may be 98.5%-100% pure is a high dose drug capable of being directly compressed with specific excipients into tablets having desired hardness, disintegrating ability, and acceptable dissoln. characteristics. Metformin HCl is not inherently compressible and thus presents formulation problems. Excipients used in the formulation enhance the flow and compaction properties of the drug and tableting mix. Optimal flow contributes to uniform die fill and wt. control. The binder used ensures sufficient cohesive properties that allow metformin HCl to be compressed using the direct compression method. The tablets produced provide an acceptable in-vitro dissoln. profile. A directly compressed tablet contained metformin HCl 500, microcryst. cellulose 36.85, hydroxypropyl Me cellulose 77.9, Povidone 26.8, colloidal silica 3.25, and Mg stearate 5.2 mg.

REFERENCE COUNT:

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT L19 ANSWER 13 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:866782 HCAPLUS

DOCUMENT NUMBER: 137:358144

Fast-release tablets containing TITLE:

metformin hydrochloride

INVENTOR(S): Matsui, Tadashi; Yuasa, Shuichiro

Toa Eiyo, Ltd., Japan PATENT ASSIGNEE(S):

Jpn. Kokai Tokkyo Koho, 5 pp. SOURCE:

CODEN: JKXXAF

DOCUMENT TYPE:

Patent Japanese LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ______ -----JP 2001-136873 20010508 JP 2002326927 A2 20021115 JP 2001-136873 20010508 PRIORITY APPLN. INFO.:

The title tablets comprise (1) 85-97.5 % metformin hydrochloride (I) and (2) 2-10 % hydroxypropyl cellulose which shows 2-10 mPas viscosity as a 2 % aq. soln. at 20.degree.. The tablets release .gtoreq. 85 % I in 15 min when tested according to Japanese Pharmacopeia XIII dissoln. test method. For example, a tablet contained I 250, hydroxypropyl cellulose (HPC SSL) 17.3, talc 1.35, and Mg stearate 1.35 mg.

L19 ANSWER 14 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:126046 USPATFULL

TITLE: Controlled release oral tablet having a

unitary core

INVENTOR(S): Cheng, Xiu Xiu, Davie, FL, UNITED STATES

Chen, Chih-Ming, Davie, FL, UNITED STATES
Jan, Steve, Coral Springs, FL, UNITED STATES
Chou, Joseph, Coral Springs, FL, UNITED STATES

APPLICATION INFO.: US 2001-16556 A1 20011101 (10)

RELATED APPLN. INFO.: Continuation of Ser. No. US 2000-594637, filed on 15

Jun 2000, PENDING

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Martin P. Endres, Esq., HEDMAN & COSTIGAN, PC., 1185

Avenue of the Americas, New York, NY, 10036

NUMBER OF CLAIMS: 30 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Page(s)

LINE COUNT: 609

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A controlled release antihyperglycemic **tablet** that does not contain an expanding polymer and comprising a core containing the antihyperglycemic drug, a semipermeable membrane **coating** the

core and at least one passageway in the membrane.

L19 ANSWER 15 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2000:101892 USPATFULL

TITLE: Controlled release oral tablet having a

unitary core

INVENTOR(S): Cheng, Xiu Xiu, Davie, FL, United States

Chen, Chih-Ming, Davie, FL, United States Jan, Steve, Coral Springs, FL, United States Chou, Joseph, Coral Springs, FL, United States

(9)

PATENT ASSIGNEE(S): Andrx Pharmaceuticals, Inc., Fort Lauderdale, FL,

United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6099859 20000808

APPLICATION INFO.: US 1998-45330 19980320

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Page, Thurman K.

ASSISTANT EXAMINER: Benston, Jr., William E.

LEGAL REPRESENTATIVE: Hedman, Gibson & Costigan, P.C.

NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 628

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A controlled release antihyperglycemic tablet that does not contain an expanding polymer and comprising a core containing the antihyperglycemic drug, a semipermeable membrane coating the

core and at least one passageway in the membrane.

L19 ANSWER 16 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2001:380381 HCAPLUS

DOCUMENT NUMBER: 134:371803

TITLE: Antidiabetic compositions containing thiazolidinedione

derivatives and metformin

INVENTOR(S): Lewis, Karen; Lillott, Nicola Jayne; Mackenzie, Donald

Colin

PATENT ASSIGNEE(S): Smithkline Beecham P.L.C., UK

SOURCE: PCT Int. Appl., 10 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO.
                                      APPLICATION NO. DATE
                  KIND DATE
    _____
                         _____
                                        _____
                          20010525
                                       WO 2000-GB4363 20001116
    WO 2001035940
                    A2
    WO 2001035940
                    A3
                          20020321
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
            HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
            LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
            SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
            YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
            BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                    A2 20020821
                                      EP 2000-976151 20001116
    EP 1231917
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
    JP 2003514011
                    T2 20030415
                                       JP 2001-537933
                                                        20001116
PRIORITY APPLN. INFO.:
                                     GB 1999-27121 A 19991116
                                     GB 2000-13238
                                                    A 20000531
                                     WO 2000-GB4363
                                                   W 20001116
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AB A pharmaceutical compn. comprises a thiazolidinedione, metformin .cntdot.HCl, and a pharmaceutically acceptable carrier, wherein the thiazolidinedione is formulated upon the surface of the metformin .cntdot.HCl. A tablet was formulated contg. metformin .cntdot.HCl 500, PVP 15, and Mg stearate 5 mg. A film coated tablet contained the above tablet 520, Opadry barrier coat 5.20, Opadry coating suspension contg. 5-[[4-[2-(methyl-2-pyridinylamino)ethoxy]phenyl]methyl]-2,4-thiazolidinedione (I) 15.90 (equiv. to 4 mg I), and Opadry I seal coat 10.80 mg.

L19 ANSWER 17 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2000:547478 HCAPLUS

DOCUMENT NUMBER: 133:155443

TITLE: Metformin formulations and method for

treating intermittent claudication employing same

INVENTOR(S): Rogosky, Karen M.

PATENT ASSIGNEE(S): Bristol-Myers Squibb Company, USA

SOURCE: U.S., 4 pp.
CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
US 6100300 A 20000808 US 1998-67565 19980428
PRIORITY APPLN. INFO.: US 1998-67565 19980428

AB Novel metformin formulations are provided which include

metformin or metformin salts preferably the

hydrochloride salt in doses below that employed for treating diabetes such as metformin in daily amts. of 400 mg or below. A method for treating peripheral vascular disease including intermittent claudication employing such metformin formulations is also provided. A tablet contained metformin.cntdot.HCl 50, microcryst. cellulose 8, Na croscarmellose 4.5, Povidone 1.5, and Mg stearate 0.8 mg.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

CCESSION NUMBER:

CORPORATE SOURCE:

1998:628926 HCAPLUS

DOCUMENT NUMBER:

130:57084

TITLE:

Improvement of quality of metformin

hydrochloride tablets by superdisintegrants

AUTHOR(S):

Wang, Xueliang; Fang, Xiaoling; Yang, Min; Zhang, Jin

Shanghai Sifu Pharmaceutical Company, Shanghai,

201106, Peop. Rep. China

SOURCE:

Zhongguo Yaoxue Zazhi (Beijing) (1998), 33(7), 434-435

CODEN: ZYZAEU; ISSN: 1001-2494

PUBLISHER:

Zhongquo Yaoxuehui

DOCUMENT TYPE:

Journal

LANGUAGE:

NGUAGE: Chinese
The improvement of quality of metformin hydrochloride

tablets in different formulations was studied. Six formulations of metformin hydrochloride tablets were designed and prepd. with microcryst. cellulose, L-HPC and three superdisintegrants (crosslinking PVP, crosslinking CMC-Na, CMS-Na) resp. The disintegration time, dissoln. rate, hardness (crushing strength) of tablets, and the granules properties were detd. and compared. The hardness of the 6 formulations were all greater than 8 kg. The disintegration time of the tablets contg. cross-linking PVP and crosslinking CMC-Na resp. were shorter than 3 min with dissoln. within 5 min. 95% Of drugs released from formulations contg. L-HPC and microcryst. cellulose within 10 min. The quality of metformin hydrochloride tablets might be significantly improved by using 4% of superdisintegrants, such as crosslinking PVP, crosslinking CMC-Na, L-HPC and CMS-Na.

L19 ANSWER 22 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN 2003:376620 HCAPLUS ACCESSION NUMBER: / DOCUMENT NUMBER: 138:374198 Controlled-release metformin tablets TITLE: Chawla, Manish; Raghuvanshi, Rajeev S.; Rampal, Ashok INVENTOR(S): PATENT ASSIGNEE(S): Ranbaxy Laboratories Limited, India PCT Int. Appl., 15 pp. SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE ______ ______ A1 20030515 WO 2002-IB4647 20021106 WO 2003039527 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG 20030605 US 2002-289070 20021106 US 2003104059 A1 IN 2001-DE1134 A 20011106 PRIORITY APPLN. INFO.: Controlled-release metformin tablets were prepd. using a combination of nonionic and anionic hydrophilic polymers, wherein the total hydrophilic polymer concn. is at least about 16% by wt. of the compn. Tablets were prepd. contg. metformin-HCl 68.0, Na CM-cellulose 4.0, HPMC 12.0, binder 1.6, diluent 13.2,

lubricant 0.6, and glidant 0.6 % wt./wt.

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

2000:68154 HCAPLUS ACCESSION NUMBER: DOCUMENT NUMBER: 132:113105 Tablets comprising a combination of TITLE: metformin and glibenclamide Bonhomme, Yves; Nicholson, Geoffrey INVENTOR(S): LIPHA, Lyonnaise Industrielle Pharmaceutique, Fr. PATENT ASSIGNEE(S): Eur. Pat. Appl., 8 pp. SOURCE: CODEN: EPXXDW DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE ---------20000126 EP 1998-401781 19980715 EP 974356 A1 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO 20000127 CA 1999-2303537 19990712 CA 2303537 AAWO 2000003742 A2 20000127 WO 1999-EP5571 20000420 WO 2000003742 **A3** AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG AU 9954179 A1 20000207 AU 1999-54179 19990712 AU 753604 B2 20021024 EP 1999-940114 19990712 EP 1011684 **A**2 20000628 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO BR 9906600 20000718 BR 1999-6600 19990712 Α **T**2 JP 2002520371 20020709 JP 2000-559876 19990712 NZ 503248 20020927 NZ 1999-503248 A 19990712 US 1999-353141 US 6303146 B1 20011016 19990714 ZA 2000-1159 ZA 2000001159 20000307 A 20010531 PRIORITY APPLN. INFO.: EP 1998-401781 A 19980715 W 19990712 WO 1999-EP5571 The present invention relates to a tablet comprising a AB

combination of metformin and glibenclamide in which the size of the glibenclamide is such that at most 10 % of the particles are less than 2 .mu.m and at most 10% of the particles are greater than 60 .mu.m. The selection of a specific size fraction of glibenclamide enables the prodn. of a combination tablet exhibiting comparable glibenclamide bioavailability to the co-administered tablets, when judged by the AUC in vivo anal. PVP 66.6 g, metformin.cntdot.HCl 1500 g, glibenclamide (10-90 % size range 2-60 .mu.m) 7.5 g, croscarmellose Na 42 g, microcryst. cellulose 284.4 g, and water 246 g were mixed and granulated. The granules were extruded through a 1 mm mesh and further mixed with microcryst. cellulose and Mg stearate. The granule mix was compressed to tablets, which were coated with a 2 % hydroxypropyl Me cellulose.

REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 24 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2000:280470 HCAPLUS

DOCUMENT NUMBER: 133:168245

TITLE: Study on HPMC matrix tablets of

metformin hydrochloride

AUTHOR(S): Xu, Qun-Wei; Hao, Qing; Zhu, Xiang-Jun

CORPORATE SOURCE: Jiangsu Institute of Materia Medica, Nanjing, 210009,

Peop. Rep. China

SOURCE: Zhongguo Yaoke Daxue Xuebao (2000), 31(1), 15-17

CODEN: ZHYXE9; ISSN: 1000-5048

PUBLISHER: Zhongguo Yaoke Daxue

DOCUMENT TYPE: Journal LANGUAGE: Chinese

The HPMC matrix tablets of metformin hydrochloride

(MH) were compressed by using wet method. The effect of the amt.,

viscosity of hydroxypropyl methylcellulose and species of bonding agent

such as Et cellulose, alc., Eudragit III on the MH release rate

from matrix tablets was investigated. The exptl. design using

orthogonal table has shown that the amt. and species of bonding agent were

affected in the MH release rate from matrix tablets and the

viscosity of HPMC was not significant.

L19 ANSWER 21 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1998:628926 HCAPLUS

DOCUMENT NUMBER: 130:57084

Improvement of quality of metformin TITLE:

hydrochloride tablets by superdisintegrants

Wang, Xueliang; Fang, Xiaoling; Yang, Min; Zhang, Jin AUTHOR(S):

Shanghai Sifu Pharmaceutical Company, Shanghai, CORPORATE SOURCE:

201106, Peop. Rep. China

Zhongquo Yaoxue Zazhi (Beijing) (1998), 33(7), 434-435 SOURCE:

CODEN: ZYZAEU; ISSN: 1001-2494

PUBLISHER: Zhongguo Yaoxuehui

DOCUMENT TYPE: Journal LANGUAGE: Chinese

The improvement of quality of metformin hydrochloride

tablets in different formulations was studied. Six formulations of metformin hydrochloride tablets were designed and prepd. with microcryst. cellulose, L-HPC and three superdisintegrants (crosslinking PVP, crosslinking CMC-Na, CMS-Na) resp. The disintegration time, dissoln. rate, hardness (crushing strength) of tablets, and the granules properties were detd. and compared. hardness of the 6 formulations were all greater than 8 kg. The disintegration time of the tablets contq. cross-linking PVP and crosslinking CMC-Na resp. were shorter than 3 min with dissoln. within 5 min. 95% Of drugs released from formulations contg. L-HPC and microcryst. cellulose within 10 min. The quality of metformin

hydrochloride tablets might be significantly improved by using 4% of superdisintegrants, such as crosslinking PVP, crosslinking CMC-Na, L-HPC and CMS-Na.

L19 ANSWER 22 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2003:376620 HCAPLUS

DOCUMENT NUMBER: 138:374198

Controlled-release metformin tablets TITLE:

Chawla, Manish; Raghuvanshi, Rajeev S.; Rampal, Ashok INVENTOR(S):

Ranbaxy Laboratories Limited, India PATENT ASSIGNEE(S):

SOURCE: PCT Int. Appl., 15 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
              KIND DATE
                                  APPLICATION NO. DATE
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                                  ______
               A1 20030515
                                  WO 2002-IB4647 20021106
WO 2003039527
   W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
       CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
       GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
       LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
       PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
       UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD,
       RU, TJ, TM
   RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
       CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
       PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
       NE, SN, TD, TG
                     20030605
                                   US 2002-289070 20021106
US 2003104059
                A1
                                IN 2001-DE1134 A 20011106
```

PRIORITY APPLN. INFO.: Controlled-release metformin tablets were prepd. using

a combination of nonionic and anionic hydrophilic polymers, wherein the total hydrophilic polymer concn. is at least about 16% by wt. of the compn. Tablets were prepd. contg. metformin-HCl 68.0, Na CM-cellulose 4.0, HPMC 12.0, binder 1.6, diluent 13.2, lubricant 0.6, and glidant 0.6 % wt./wt.

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 23 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2000:68154 HCAPLUS

DOCUMENT NUMBER: 132:113105

Tablets comprising a combination of TITLE:

metformin and glibenclamide

INVENTOR(S): Bonhomme, Yves; Nicholson, Geoffrey

LIPHA, Lyonnaise Industrielle Pharmaceutique, Fr. PATENT ASSIGNEE(S):

Eur. Pat. Appl., 8 pp. SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO. · KIND DATE
                                 APPLICATION NO. DATE
    EP 974356 A1 20000126 EP 1998-401781 19980715
    -----
                                       ______
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
           IE, SI, LT, LV, FI, RO
                   AA 20000127
                                       CA 1999-2303537 19990712
    CA 2303537
                                      WO 1999-EP5571 19990712
    WO 2000003742
                    A2
                         20000127
    WO 2000003742
                   A3
                        20000420
           AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,
           DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
           JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
           MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
           TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
           MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
           ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
           CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                   A1
                                      AU 1999-54179
    AU 9954179
                         20000207
                                                     19990712
    AU 753604
                    B2
                         20021024
                                       EP 1999-940114 19990712
    EP 1011684
                    A2
                         20000628
          AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
           IE, SI, LT, LV, FI, RO
                                       BR 1999-6600
                                                      19990712
    BR 9906600
                   Α
                        20000718
    JP 2002520371
                    T2
                         20020709
                                      JP 2000-559876 19990712
    NZ 503248
                    Α
                         20020927
                                      NZ 1999-503248 19990712
    US 6303146
                    В1
                         20011016
                                       US 1999-353141 19990714
    ZA 2000001159
                   Α
                        20010531
                                       ZA 2000-1159
                                                      20000307
PRIORITY APPLN. INFO.:
                                    EP 1998-401781 A 19980715
                                    WO 1999-EP5571 W 19990712
    The present invention relates to a tablet comprising a
AB
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combination of metformin and glibenclamide in which the size of the glibenclamide is such that at most 10 % of the particles are less than 2 .mu.m and at most 10% of the particles are greater than 60 .mu.m. The selection of a specific size fraction of glibenclamide enables the prodn. of a combination tablet exhibiting comparable glibenclamide bioavailability to the co-administered tablets, when judged by the AUC in vivo anal. PVP 66.6 g, metformin.cntdot.HCl 1500 g, glibenclamide (10-90 % size range 2-60 .mu.m) 7.5 g, croscarmellose Na 42 g, microcryst. cellulose 284.4 g, and water 246 g were mixed and granulated. The granules were extruded through a 1 mm mesh and further mixed with microcryst. cellulose and Mg stearate. The granule mix was compressed to tablets, which were coated with a 2 % hydroxypropyl Me cellulose.

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 2 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT ACCESSION NUMBER: 2000:280470 HCAPLUS

DOCUMENT NUMBER: 133:168245

TITLE: Study on HPMC matrix tablets of

metformin hydrochloride

AUTHOR(S): Xu, Qun-Wei; Hao, Qing; Zhu, Xiang-Jun

CORPORATE SOURCE: Jiangsu Institute of Materia Medica, Nanjing, 210009,

Peop. Rep. China

SOURCE: Zhongquo Yaoke Daxue Xuebao (2000), 31(1), 15-17

CODEN: ZHYXE9; ISSN: 1000-5048

PUBLISHER: Zhongguo Yaoke Daxue

DOCUMENT TYPE: Journal LANGUAGE: Chinese

AB The HPMC matrix tablets of metformin hydrochloride

(MH) were compressed by using wet method. The effect of the amt., viscosity of hydroxypropyl methylcellulose and species of bonding agent such as Et cellulose, alc., Eudragit III on the MH release rate from matrix tablets was investigated. The exptl. design using orthogonal table has shown that the amt. and species of bonding agent were affected in the MH release rate from matrix tablets and the

viscosity of HPMC was not significant.

L19 ANSWER 25 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:72923 USPATFULL

TITLE: Liquid formulation of metformin

INVENTOR(S): Chandran, Ravi, Bolton Landing, NY, UNITED STATES

Gogia, Ashish, New Delhi, INDIA

NUMBER DATE

PRIORITY INFORMATION: US 2000-223391P 20000807 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: RANBAXY PHARMACEUTICALS INC., Suite 2100, 600 College

Road East, Princeton, NJ, 08540

NUMBER OF CLAIMS: 50 EXEMPLARY CLAIM: 1 LINE COUNT: 1042

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention is directed to a liquid formulation of metformin or its pharmaceutically acceptable salts thereof. The liquid pharmaceutical composition comprises a therapeutically effective amount of metformin or its pharmaceutically acceptable salt, in a liquid carrier, which may also include a sweetener that does not increase the blood glucose level of a subject after ingestion thereof. In one embodiment, it may also include alkyl hydroxyethylcellulose, and/or a polyhydroxy alcohol. In another embodiment, the carrier may contain a sweetener, mineral acid, and bicarbonate salt maintained at a pH of 4.0 to 9.0. It is useful for treating hyperglycemia and diabetes.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 26 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:214468 USPATFULL

TITLE: Liquid formulation of metformin

INVENTOR(S): Chandran, Ravi, Bolton Landing, NY, UNITED STATES

Gogia, Ashish, New Delhi, INDIA

NUMBER KIND DATE

US 2003149111 A1 20030807 US 2003-382442 A1 20030306 (10) PATENT INFORMATION:

APPLICATION INFO.:

Continuation of Ser. No. US 2001-923491, filed on 7 Aug RELATED APPLN. INFO.:

2001, GRANTED, Pat. No. US 6559187

NUMBER DATE

------PRIORITY INFORMATION: US 2000-223391P 20000807 (60)

DOCUMENT TYPE: Utility

APPLICATION FILE SEGMENT:

JAYADEEP R. DESHMUKH, ESQ., RANBAXY PHARMACEUTICALS LEGAL REPRESENTATIVE:

INC., Suite 2100, 600 College Road East, Princeton, NJ,

50 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1044 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention is directed to a liquid formulation of

metformin or its pharmaceutically acceptable salts thereof. The liquid pharmaceutical composition comprises a therapeutically effective

amount of metformin or its pharmaceutically acceptable salt,

in a liquid carrier, which may also include a sweetener that does not increase the blood glucose level of a subject after ingestion thereof. In one embodiment, it may also include alkyl hydroxyethylcellulose, and/or a polyhydroxy alcohol. In another embodiment, the carrier may contain a sweetener, mineral acid, and bicarbonate salt maintained at a pH of 4.0 to 9.0. It is useful for treating hyperglycemia and diabetes.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 27 OF 256 USPATFULL on STN

2001:.147493 USPATFULL ACCESSION NUMBER:

Controlled release tablet having a unitary TITLE:

Chen, Chih-Ming, Davie, FL, United States INVENTOR(S):

Cheng, Xiu Xiu, Davie, FL, United States

Chou, Joseph, Coral Springs, FL, United States Jan, Steve, Coral Springs, FL, United States

Andrx Pharmaceuticals, Inc., Davie, FL, United States PATENT ASSIGNEE(S):

(U.S. corporation)

NUMBER KIND DATE _____

US 6284275 B1 20010904 PATENT INFORMATION: APPLICATION INFO.: US 2000-590807 20000609 (9)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1998-143876, filed on 31

Aug 1998, now patented, Pat. No. US 6099862

DOCUMENT TYPE: Utility GRANTED FILE SEGMENT:

Page, Thurman K. PRIMARY EXAMINER: Seidleck, Brian K. ASSISTANT EXAMINER: LEGAL REPRESENTATIVE: Hedman & Costigan P.C.

NUMBER OF CLAIMS: 39 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 2 Drawing Figure(s); 2 Drawing Page(s)

LINE COUNT: 639

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A controlled release pharmaceutical tablet containing AB

antihyperglycemic drug and a hypoglycemic drug that does not contain an expanding or gelling polymer layer and comprising a core containing the antihyperglycemic drug and the hypoglycemic drug, a semipermeable

coating membrane surrounding the core and at least one

passageway in the membrane to allow the drugs to be released from the

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 28 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2003:282377 HCAPLUS

DOCUMENT NUMBER: 138:292793

Extended release pharmaceutical composition containing TITLE:

metformin

Murpani, Deepak; Madan, Ashish; Arora, Vinod Kumar; INVENTOR(S):

Malik, Rajiv

Ranbaxy Laboratories Limited, India PATENT ASSIGNEE(S):

SOURCE: PCT Int. Appl., 18 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
              KIND DATE
                                  APPLICATION NO. DATE
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                                   -----
WO 2003028704
               A1
                     20030410
                                   WO 2002-IB3997
                                                  20020927
   W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
       CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
       GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
       LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
       PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
       UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD,
       RU, TJ, TM
   RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
       CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
       PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
       NE, SN, TD, TG
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PRIORITY APPLN. INFO.:

IN 2001-DE1002 A 20010928

The present invention relates to an extended release pharmaceutical compn. contg. metformin and a rate controlling polymer and a process for its prepn. are described. The compn. has a water content of 3.2-10.0% by wt. and improved hardness and friability. For example, tablets with water content of 2.8% were prepd. by conventional dry granulation technique from a blend of metformin hydrochloride 500.0 mg, sodium CM-cellulose 36.0 mg, microcryst. cellulose 60.0 mg, hydroxypropyl Me cellulose 398.0 mg, magnesium stearate 6 mg, and water as needed. Hardness of the tablets obtained was 16.9 Kp and friability was 0.43% by wt. Release of metformin hydrochloride from tablets after 1h, 4 h, 8 h, and 12 h was

27.1%, 58.7%, 84.9%, and 97.8%, resp.

REFERENCE COUNT: THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 29 OF 256 USPATFULL on STN

2001:25927 USPATFULL ACCESSION NUMBER:

TITLE: Method of reducing serum glucose levels

INVENTOR (S): Byrd, Edward A., San Francisco, CA, United States

Janjikhel, Rajiv, Owings Mills, MD, United States

Medical Research Institute, San Bruno, CA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

	NUMBER	KIND	DATE	
		- -		
PATENT INFORMATION:	US 6191162	B1	20010220	
APPLICATION INFO.:	US 1999-288253		19990408	(9

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1998-112623, filed

on 9 Jul 1998

NUMBER DATE ______

US 1998-102605P 19981001 (60) PRIORITY INFORMATION:

US 1998-87203P 19980528 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Criares, Theodore J. PRIMARY EXAMINER:

ASSISTANT EXAMINER: Kim, Jennifer

Bozicevic, KarlBozicevic, Field, Francis LLP LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1694 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A controlled release formulation of lipoic acid is administered to a patient resulting in reduced serum glucose levels. The formulation comprises a pharmaceutically acceptable carrier and is designed to prevent degradation of the lipoic acid in the gastrointestinal tract and to release the lipoic acid in a controlled manner thereby obtaining a desired lipoic acid serum level over an extended period resulting in reduced serum glucose levels over that period.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 30 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2001:144935 USPATFULL

EXTENDING THE DURATION OF DRUG RELEASE WITHIN THE TITLE:

STOMACH DURING THE FED MODE

SHELL, JOHN W., HILLSBOROUGH, CA, United States INVENTOR(S):

LOUIE-HELM, JENNY, UNION CITY, CA, United States MARKEY, MICHELINE, SANTA CRUZ, CA, United States

NUMBER KIND DATE -----US 2001018070 A1 20010830 US 6340475 B2 20020122 US 1999-282233 A1 19990329 (9) PATENT INFORMATION: APPLICATION INFO.:

Continuation-in-part of Ser. No. US 1997-870509, filed RELATED APPLN. INFO.:

on 6 Jun 1997, ABANDONED A 371 of International Ser.

No. WO 1998-US11302, filed on 5 Jun 1998, UNKNOWN

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

M HENRY HEINES, TOWNSEND TOWNSEND & CREW, TWO LEGAL REPRESENTATIVE:

EMBARCADERO CENTER, 8TH FLOOR, SAN FRANCISCO, CA,

941113834

NUMBER OF CLAIMS: 44 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 9 Drawing Page(s)

LINE COUNT: 1530

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Drugs are formulated as unit oral dosage forms by incorporating them into polymeric matrices comprised of hydrophilic polymers that swell upon imbibition of water to a size that is large enough to promote retention of the dosage form in the stomach during the fed mode. The oral formulation is designed for gastric retention and controlled delivery of an incorporated drug into the gastric cavity, and thus administered, the drug is released from the matrix into the gastric fluid by solution diffusion. The swollen polymeric matrix, having achieved sufficient size, remains in the gastric cavity for several hours if administered while the patient is in the fed mode, and remains intact long enough for substantially all of the drug to be released before substantial dissolution of the matrix occurs. The swelling matrix lowers the accessibility of the gastric fluid to the drug and thereby reduces the drug release rate. This process, together with diffusion retardation by selection of specific polymers, polymer molecular

weights, and other variables, results in a sustained and controlled delivery rate of the drug to the gastric cavity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 31 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:98925 USPATFULL

TITLE: Extending the duration of drug release within the

stomach during the fed mode

INVENTOR(S): Shell, John W., Hillsborough, CA, UNITED STATES

Louie-Helm, Jenny, Union City, CA, UNITED STATES Markey, Micheline, Santa Cruz, CA, UNITED STATES

PATENT ASSIGNEE(S): DepoMed, Inc., Menlo Park, CA, UNITED STATES (U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 2002051820 A1 20020502 APPLICATION INFO.: US 2001-990061 A1 20011120 (9)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1999-282233, filed on 29

Mar 1999, PENDING Continuation-in-part of Ser. No. US

1997-870509, filed on 6 Jun 1997, ABANDONED

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO

CENTER, EIGHTH FLOOR, SAN FRANCISCO, CA, 94111-3834

NUMBER OF CLAIMS: 44 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 9 Drawing Page(s)

LINE COUNT: 1493

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Drugs are formulated as unit oral dosage forms by incorporating them into polymeric matrices comprised of hydrophilic polymers that swell upon imbibition of water to a size that is large enough to promote retention of the dosage form in the stomach during the fed mode. The oral formulation is designed for gastric retention and controlled delivery of an incorporated drug into the gastric cavity, and thus administered, the drug is released from the matrix into the gastric fluid by solution diffusion. The swollen polymeric matrix, having achieved sufficient size, remains in the gastric cavity for several hours if administered while the patient is in the fed mode, and remains intact long enough for substantially all of the drug to be released before substantial dissolution of the matrix occurs. The swelling matrix lowers the accessibility of the gastric fluid to the drug and thereby reduces the drug release rate. This process, together with diffusion retardation by selection of specific polymers, polymer molecular weights, and other variables, results in a sustained and controlled delivery rate of the drug to the gastric cavity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 32 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:57124 USPATFULL

TITLE: Extending the duration of drug release within the

stomach during the fed mode

INVENTOR(S): Shell, John W., Hillsborough, CA, UNITED STATES

Louie-Helm, Jenny, Union City, CA, UNITED STATES Markey, Micheline, Santa Cruz, CA, UNITED STATES

PATENT ASSIGNEE(S): DepoMed, Inc., Menlo Park, CA, 94025 (U.S. corporation)

Continuation of Ser. No. US 1999-282233, filed on 29 RELATED APPLN. INFO.:

Mar 1999, GRANTED, Pat. No. US 6340475

Continuation-in-part of Ser. No. US 1997-870509, filed

on 6 Jun 1997, ABANDONED

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO

CENTER, EIGHTH FLOOR, SAN FRANCISCO, CA, 94111-3834

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

9 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 1439

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Drugs are formulated as unit oral dosage forms by incorporating them into polymeric matrices comprised of hydrophilic polymers that swell upon imbibition of water to a size that is large enough to promote retention of the dosage form in the stomach during the fed mode. The oral formulation is designed for gastric retention and controlled delivery of an incorporated drug into the gastric cavity, and thus administered, the drug is released from the matrix into the gastric fluid by solution diffusion. The swollen polymeric matrix, having achieved sufficient size, remains in the gastric cavity for several hours if administered while the patient is in the fed mode, and remains intact long enough for substantially all of the drug to be released before substantial dissolution of the matrix occurs. The swelling matrix lowers the accessibility of the gastric fluid to the drug and thereby reduces the drug release rate. This process, together with diffusion retardation by selection of specific polymers, polymer molecular weights, and other variables, results in a sustained and controlled delivery rate of the drug to the gastric cavity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 33 OF 256 USPATFULL on STN

2003:133545 USPATFULL ACCESSION NUMBER:

TITLE:

Formulation of an erodible, gastric retentive oral dosage form using in vitro disintegration test data

A1 20011025 (10)

Louie-Helm, Jenny, Union City, CA, UNITED STATES

Berner, Bret, El Granada, CA, UNITED STATES

NUMBER KIND DATE US 2003091630 A1 20030515 PATENT INFORMATION:

US 2001-14750 APPLICATION INFO.: DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

REED & EBERLE LLP, 800 MENLO AVENUE, SUITE 210, MENLO LEGAL REPRESENTATIVE:

PARK, CA, 94025

NUMBER OF CLAIMS: 44 EXEMPLARY CLAIM:

INVENTOR(S):

NUMBER OF DRAWINGS: 5 Drawing Page(s)

LINE COUNT: 1906

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Erodible, gastric-retentive dosage forms are provided that are AB formulated using the in vitro drug release profile obtained with USP Disintegration test equipment rather the USP Dissolution Apparatus. The invention is premised on the discovery that the USP Disintegration Test and modified versions thereof are far more predictive of the in vivo release profile for a controlled release dosage form than is the standard USP Dissolution Test, particularly controlled release dosage forms of the swellable, erodible type. The dosage forms generally comprise particles of a biocompatible, hydrophilic polymer having the active agent incorporated therein, wherein the particles are optionally but preferably compacted into a tablet or loaded into a

capsule. The dosage forms can be used to deliver water-insoluble or sparingly soluble drugs as well as water-soluble drugs, providing that the latter are coated with a protective coating or contained in a protective vesicle.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 34 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1997:128753 HCAPLUS

DOCUMENT NUMBER: 126:229547

Use of cellulose ether containing excipients TITLE:

> with microcrystalline cellulose for the production of pellets containing metformin hydrochloride by the process of extrusion-

spheronization

Gouldson, M. P.; Deasy, P. B. AUTHOR (S):

Dep. Pharmaceutics, Trinity Coll. Univ. Dublin, CORPORATE SOURCE:

Dublin, 4, Ire.

Journal of Microencapsulation (1997), 14(2), 137-153 SOURCE:

CODEN: JOMIEF; ISSN: 0265-2048

Taylor & Francis PUBLISHER:

DOCUMENT TYPE: Journal LANGUAGE: English

The project is concerned mainly with the evaluation of 2 cellulose ether contg. excipients, Aquacoat WG and Avicel 955 MCC for the improved

extrusion-spheronization of metformin-HCl. Factorially designed expts. subject to statistical analyses were employed and products obtained

were evaluated by sieve, packing d. and image anal., SEM and dissoln. testing at pH 6.cntdot.8. Aquacoat WG did not improve the ease of spheronization of drug mixes contg. microcryst. cellulose wetted with the optimum level of water. However, Avicel 955 MCC, a new exptl.

excipient contg. 95% microcryst. cellulose and 5% Me

cellulose, did aid ease of spheronization facilitating acceptable yield of good spheres with high drug loadings (70%). Avicel 955

MCC-contq. drug mixes were more tolerant to minor alterations in level of hydration and yielded spheres which showed a small retardation of drug release despite the very high soly. of metformin-HCl.

L19 ANSWER 35 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2001:178655 USPATFULL

TITLE: Solid oral dosage form comprising a combination of

metformin and glibenclamide

INVENTOR(S): Bonhomme, Yves, Charbonnieres les Bains, France

Nicholson, Geoffrey, Aylesbury, United Kingdom Cave, Gillian, Ellesmere Port, United Kingdom Nicholson, Sarah J., Helsby, United Kingdom

PATENT ASSIGNEE(S): LIPHA, Lyons, France (non-U.S. corporation)

NUMBER KIND DATE ---**--**----US 6303146 B1 20011016 PATENT INFORMATION: APPLICATION INFO.: US 1999-353141 19990714 (9)

NUMBER DATE -----

PRIORITY INFORMATION: EP 1998-401781 19980715

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

Page, Thurman K. PRIMARY EXAMINER:

ASSISTANT EXAMINER: Tran, S.

Oblon, Spivak, McClelland, Maier & Neustadt, P.C. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 3 Drawing Figure(s); 3 Drawing Page(s) LINE COUNT: 418

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to a solid oral dosage form comprising a combination of metformin and glibenclamide in which the size of glibenclamide is such that the glibenclamide bioavailability is comparable to the glibenclamide bioavailability obtained with a separate administration of metformin and glibenclamide.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 36 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2003:150455 HCAPLUS

DOCUMENT NUMBER: 138:175909

Directly compressible extended-release matrix TITLE:

formulation for metformin hydrochloride

Kumar, Vijai; McGuffy, Kevin Scott INVENTOR(S):

PATENT ASSIGNEE(S): USA

U.S., 7 pp. SOURCE:

CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ---------B1 20030225 US 2001-8/9/40 20010612
US 2001-879748 20010612 US 6524618 PRIORITY APPLN. INFO.: An extended-release matrix formulation capable of being directly compressed into tablets comprises metformin-HCl blended with specific excipients. The excipients used in the formulation enhance the flow and compaction properties of the drug and insure that the formulation is directly compressible into a tablet contg. 100-800 mg, preferably 250-750 mg, of metformin-HCl in unit dosage form. Each tablet produced by direct compression of the formulation has the desired hardness and dissoln. characteristics such that the drug is released in the body of the subject over an extended period of time. Tablets were prepd. from metformin -HCl 750.00, lactose 161.55, hydroxypropyl cellulose 463.50,

hydroxyethyl cellulose 154.50, colloidal silicon dioxide 7.73, and Mg stearate 7.72 mg.

THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 14 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 37 OF 256 HCAPLUS. COPYRIGHT 2003 ACS on STN

2001:899701 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 136:74508

In vitro comparative study of biopharmaceutical TITLE:

properties of metformin hydrochloride

tablets marketed in Brazil

Gomes de Pinho, Jose de Jesus Ribeiro; Storpirtis, AUTHOR (S):

Silvia

Fac. Farmacia Bioquimica, Univ. Federal Juiz de Fora, CORPORATE SOURCE:

Brazil

Revista Brasileira de Ciencias Farmaceuticas (2001), SOURCE:

37(1), 95-105

CODEN: RBCFFM; ISSN: 1516-9332

Universidade de Sao Paulo, Faculdade de Ciencias PUBLISHER:

Farmaceuticas

DOCUMENT TYPE: Journal LANGUAGE: Portuguese

In the present work metformin hydrocloride 850 mg

tablets, from two different labs. A and B (three batches of each lab.), were evaluated using phys. and physicochem. expts. according to British Pharmacopea (1993). The drug was assayed using UV spectrophotometry at 233 nm. The results showed that two batches from lab. B were not according to the specification because they presented irregular hardness 2.57 .+-. 0.98 and 2.89 .+-. 0.62 kgf, under minimal values of Farmacopeia Brasileira 4. ed. (Parte I), which is 3 kgf. All the batches from lab. A, which had film **coating**, showed irregular hardness (22.99 .+-. 1.49, 8.64 .+-. 0.99 and 19.02 .+-. 2.36). The products A and B developed different dissoln. profiles, resulting in order 1 kinetic. The dissoln. rate from the product A was the lowest, presenting dissoln. rate const. (Kd = 0.0518 min-1), dissoln. half-life (Td50 = 6.93 min), dissoln. efficiency (DE = 74,75%) and correlation coeff. (r = 0.9885), while the product B showed Kd = 0.0703; Td50 = 4.47 min; DE = 80,46% and r = 0.9986. Thermoanalitical tests TG/DTG and DSC demonstrated that the products suffered thermal decompn. in different temps., which can be attributed to the excipients which are distinct in the formulations.

REFERENCE COUNT:

26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 38 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2001:176241 USPATFULL

TITLE: Controlled release lipoic acid

INVENTOR(S): Byrd, Edward A., San Francisco, CA, United States

NUMBER KIND DATE

PATENT INFORMATION: US 2001028896 A1 20011011

US 6572888 B2 20030603

APPLICATION INFO.: US 2001-755890 A1 20010105 (9)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-288245, filed

on 8 Apr 1999, GRANTED, Pat. No. US 6197340

Continuation-in-part of Ser. No. US 1998-112623, filed

on 9 Jul 1998, ABANDONED

US 1998-87203P
DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Karl Bozicevic, BOZICEVIC, FIELD & FRANCIS LLP, 200

Middlefield Road, Suite 200, Menlo Park, CA, 94025

NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 1 Drawing Page(s)

LINE COUNT: 1438

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A controlled release formulation of lipoic acid is disclosed. The lipoic acid is combined with excipient materials in such a way that those materials provide for gradual release of the lipoic acid in a manner which makes it possible to substantially increase the period of time over which therapeutic levels of lipoic acid are maintained relative to a quick release formulation. These features make it possible to use lipoic acid to reduce serum glucose levels and maintain those levels over time thereby obtaining a range of desired therapeutic results.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 39 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2000:84280 HCAPLUS

DOCUMENT NUMBER: 132:127735

TITLE: Tablets for extended release of a drug in

the stomach

INVENTOR(S): Bonhomme, Yves; Nicholson, Geoffroy

LIPHA, Lyonnaise Industrielle Pharmaceutique, Fr. PATENT ASSIGNEE(S):

Eur. Pat. Appl., 8 pp. SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE EP 976395 A1 20000202 EP 1998-401956 19980730 EP 976395

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

IE, SI, LT, LV, FI, RO

AU 9957318 A1 20000221 AU 1999-57318 WO 1999-EP5746 19990729 A1 20000210 WO 2000006129

W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,

DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,

MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,

CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.:

EP 1998-401956 A 19980730 WO 1999-EP5746 W 19990729

The invention relates to a tablet for extended release of a drug ABin the stomach, comprising granules of the drug in a hydrophilic matrix, the granules being coated with a coating comprising a source of a carbon dioxide and the coating granules being blended with an agent inducing the release of carbon dioxide and tabletting aids. Granules were formulated contg. metformin.cntdot.HCl 62.42, Methocel K100M 15.9, and PVP K30 4.6 % and the granules were sprayed with PVP K30 1.6 and NaHCO3 12 % and mixed with citric acid 2.1 and Mg stearate 1.22 % for compression to give a tablet contg. metformin .cntdot.HCl 500 mg/each.

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 40 OF 256 USPATFULL on STN

3

ACCESSION NUMBER:

2003:29898 USPATFULL

TITLE: INVENTOR(S): Pharmaceutical composition Matharu, Amol Singh, Cranbury, NJ, UNITED STATES

Patel, Mahendra R., East Brunswick, NJ, UNITED STATES

KIND DATE NUMBER _______ US 2003021841 A1 US 2002-183881 A1 20030130 PATENT INFORMATION: 20020627 (10) APPLICATION INFO.:

> NUMBER DATE _____

PRIORITY INFORMATION:

US 2001-302613P 20010702 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

THOMAS HOXIE, NOVARTIS CORPORATION, PATENT AND TRADEMARK DEPT, 564 MORRIS AVENUE, SUMMIT, NJ,

079011027

NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: 1 565 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to a process for preparing tablet dosage forms of poorly-compressible pharmaceutical agents and to tablet dosage forms prepared according to the inventive process. The inventive process is especially useful for preparing tablets of the poorly-compressible drug metformin HCl.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 41 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:152383 USPATFULL

TITLE: Metformin Hydrochloride tablets

INVENTOR(S): Sherman, Bernard Charles, Willowdale, CANADA

PATENT INFORMATION: US 2003104049 A1 20030605 APPLICATION INFO.: US 2001-2130 A1 20011205 (10)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: NIXON & VANDERHYE P.C., 8th Floor, 1100 North Glebe

Road, Arlington, VA, 22201-4714

NUMBER OF CLAIMS: 20 EXEMPLARY CLAIM: 1 LINE COUNT: 209

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Tablets for oral administration comprising metformin

hydrochloride and methylcellulose.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 42 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1996:226313 HCAPLUS

DOCUMENT NUMBER: 124:270593

TITLE: Metformin controlled-release formulations

INVENTOR(S): Moeckel, Joern; Gabel, Rolf-Dieter; Woog, Heinrich

PATENT ASSIGNEE(S): boehringer Mannheim Gmbh, Germany

SOURCE: Ger. Offen., 13 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

:	PAT	ENT	NO.		KII	ND	DATE			A	PPLI	CATI	ON N	ο.	DATE			
				-						-					- -			
]	DE	4432	757		A:	1.	1996	0321		D	E 19	94-4	4327	57	1994	0914		
	ZA	9507	670		Α		1997	0313		Z	A 19	95-7	670		1995	0913		
1	WO	9608	243		A:	1.	1996	0321		W	0 19	95-E	P361	0	1995	0914		
		W:	ΑU,	ВG,	BR,	BY,	CA,	CN,	CZ,	EE,	FI,	GE,	HU,	JΡ,	∙KR,	ΚZ,	LT,	LV,
			MX,	NO,	NZ,	PL,	RO,	RU,	SI,	SK,	UA,	US						
		RW:	ΑT,	ВE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	ΝL,	PT,	SE
1	ΑU	9535	672		A:	1	1996	0329		A	U 19	95-3	5672		1995	0914		
1	ΕP	7811	29		A:	1	1997	0702		E	P 19	95-9	3274	1	1995	0914		
1	EΡ	7811	29		B	1	2003	0702										
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙE,	IT,	LI,	LU,	NL,	PT,	SE
	JΡ	1050	5604		T	2	1998	0602		J	P 19	95-5	0991	5	1995	0914		
	$_{ ext{IL}}$	1153				1	2000	0831		I:	և 19	95-1	1530	9	1995	914		
1	ΑТ	2440	04		\mathbf{E}										1995			
1	US	5955	106		A		1999	0921		U	S 19	97-7	9375	3	1997	0314		
PRIOR	ITY	APP	LN.	INFO	. :					DE 1	994 -	4432	757	Α	1994	914		
									1	WO 1	995-	EP36	10	W	19950	914		

AB Metformin is formulated with a hydrocolloid-forming substance (e.g. a gum, cellulose deriv., or synthetic polymer) as release-controlling agent with a residual moisture content of 0.5-3 wt.%. These formulations can be compressed into tablets or pellets without use of org. solvents, and can be prepd. with a high

metformin content. Thus, tablet cores were prepd. each contg. metformin-HCl 850.00, hydroxypropylmethylcellulose 60.00, PVP 38.00, and Mg stearate 5.00 mg, and coated with a mixt. of hydroxypropylmethylcellulose 20.00, ethylcellulose 12.00, Macrogol 4.00, and TiO2 4.00 mg.

L19 ANSWER 43 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:130005 USPATFULL

TITLE: Composition containing ascorbic acid INVENTOR(S): Noguchi, Hiroshi, Kawanishi, JAPAN

Taiji, Mutsuo, Takatsuki, JAPAN Yamaga, Hiroshi, Suita, JAPAN Itakura, Yasushi, Nara, JAPAN Ichihara, Junji, Takatsuki, JAPAN

PATENT ASSIGNEE(S): Sumitomo Pharmaceuticals Co., Ltd., Osaka, JAPAN

(non-U.S. corporation)

19990609 PCT 371 date

NUMBER DATE

PRIORITY INFORMATION: JP 1996-356302 19961224

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Krass, Frederick ASSISTANT EXAMINER: Jagoe, Donna

LEGAL REPRESENTATIVE: Birch, Stewart, Kolasch & Birch, LLP

NUMBER OF CLAIMS: 6 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT: 787

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L-ascorbic acid, L-ascorbic acid derivatives and salts thereof can reduce lactic acid levels in blood, and are useful for treating lactic acidosis and the like caused by administration of amoxapine, theophylline, metformin, phenformin, buformin, nalidixic acid, hopantenic acid, azidothymidine, dideoxycytidine, high caloric transfusion, propylene glycol, ethylene glycol, xylitol, lactose, sorbitol or the like.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 44 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:353262 HCAPLUS

DOCUMENT NUMBER: 136:345841

TITLE: Controlled release metformin compositions

INVENTOR(S): Chen, Chih-Ming; Cheng, Xiu-Xiu; Jan, Steve; Chou,

Joseph

PATENT ASSIGNEE(S): Andrx Corporation, USA SOURCE: PCT Int. Appl., 52 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

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C2
                           20030724
    WO 2002036100
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
            RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
            UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
            BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                      AU 2002-30830
                     A5
                          20020515
    AU 2002030830
                                         EP 2001-991078 20011030
     EP 1335708
                      A1
                          20030820
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                                       US 2000-705625
PRIORITY APPLN. INFO.:
                                                        A 20001103
                                       US 2000-705630
                                                       A 20001103
                                       WO 2001-US48306 W 20011030
AΒ
    A compn. and methods thereof for treating patients having
    non-insulin-dependent diabetes mellitus (NIDDM) by administering a
    controlled release oral solid dosage form contg. preferably a biguanide
    drug such as metformin, on a once-a-day basis. The dosage form
    provides a mean time to max. plasma concn. (Tmax) of the drug which occurs
    at 5.5 to 7.5 h after oral administration on a one-a-day basis to human
    patients. Preferably, the dose of drug is administered at dinner time to
    a patient in the fed state. A tablet core was formulated contg.
    metformin.cntdot.HCl 500, Povidone 36, Na lauryl sulfate 25.8, and
    Mg stearate 2.8 mg/tablet was coated to have a sustained-release
    coating contg. cellulose acetate 21.5, triacetin 1.3,
     and PEG-400 2.5 mg/tablet. The coated tablets were
     laser drilled two holes.
REFERENCE COUNT:
                               THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L19 ANSWER 45 OF 256 HCAPLUS- COPYRIGHT 2003 ACS on STN
                        2002:881038 HCAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                        139:57791
TITLE:
                        Preparation and in vitro release of intragastric
                        floating system of metformin hydrochloride
AUTHOR (S):
                        Huang, Dong-po; Wang, Yuan; Jiang, Guo-qiang; Chen,
                        Jun; Ding, Fu-xin
CORPORATE SOURCE:
                        Department of Chemical Engineering, Tsinghua
                        University, Beijing, 100084, Peop. Rep. China
SOURCE:
                        Jingxi Huagong (2002), 19(10), 609-611
                        CODEN: JIHUFJ; ISSN: 1003-5214
                        Jingxi Huagong Bianjibu
PUBLISHER:
DOCUMENT TYPE:
                        Journal
LANGUAGE:
                        Chinese
AB
    Intragastric floating sustained release tablets of
    metformin hydrochloride were prepd. utilizing the technique of wet
    granulation followed by compression into tablets. The
     tablets possessed superior floating property and could hold
    consistent drug release rate within over 8 h. The floating lag time
    decreased with increase in the hydroxypropyl Me cellulose
    content in the tablet. The relation between the tablet
    d. and the mass fraction of octadecyl alc. can be correlated.
    vitro release results indicated that the drug release was attributed to
    dual function of diffusion and matrix dissoln. and the kinetics was found
    to follow the Higuchi equation.
L19 ANSWER 46 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
                        2003:417505 HCAPLUS
ACCESSION NUMBER:
```

WO 2002036100

DOCUMENT NUMBER:

TITLE:

139:12256

Pharmaceutical composition containing

A1

20020510

WO 2001-US48306 20011030

metformin and a 4-oxobutanoic acid for the

treatment of diabetes

INVENTOR(S): Moinet, Gerard; Marais, Dominique

Lipha, Fr. PATENT ASSIGNEE(S):

Fr. Demande, 21 pp. SOURCE:

CODEN: FRXXBL

DOCUMENT TYPE: Patent French LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

```
APPLICATION NO. DATE
            KIND DATE
PATENT NO.
_____
                                   ______
              A1 20030530
                                FR 2001-15398 20011128
WO 2002-EP12355 20021106
FR 2832633
               A1 20030605
WO 2003045368
   W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
       CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
       GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
       LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
       PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
       UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
       TJ, TM
   RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
       CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
       PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
       NE, SN, TD, TG
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PRIORITY APPLN. INFO.:

FR 2001-15398 A 20011128

MARPAT 139:12256 OTHER SOURCE(S):

Pharmaceutical compn. comprise metformin or its pharmaceutically acceptable salts and acids and a 4-oxo-butanoic acid deriv., in combination with one or more excipients. The compns. are particularly useful for the treatment of the noninsulino-dependent diabetes. A tablet contained metformin 7.7, microcryst.

cellulose 76.7, lactose powder 4.6, hydroxy pr cellulose 1.8, sodium croscarmellose 1.8, colloidal silica (Aerosil) 0.3, and magnesium stearate 0.9%.

REFERENCE COUNT:

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 47 OF 256 USPATFULL on STN

2002:239028 USPATFULL ACCESSION NUMBER:

2

Inhibition of emetic effect of metformin with TITLE:

5-HT3 receptor antagonists

Cowles, Verne E., Dublin, CA, United States INVENTOR(S):

DepoMed, Inc., Menlo Park, CA, United States (U.S. PATENT ASSIGNEE(S):

corporation)

	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 6451808	B1	20020917	
APPLICATION INFO.:	US 2000-691398		20001017	(9)
DOCUMENT TYPE:	Utility			
FILE SEGMENT:	GRANTED			
PRIMARY EXAMINER:	Pryor, Alton Nath	aniel		

Townsend and Townsend and Crew, LLP LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: 11 EXEMPLARY CLAIM:

0 Drawing Figure(s); 0 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 444

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Metformin is formulated as a pharmaceutical composition that AB also includes a 5-hydroxytryptamine-3 receptor antagonist to suppress the gastrointestinal side effects that are associated with metformin administration in many patients.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 48 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2003:434338 HCAPLUS

DOCUMENT NUMBER: 139:12295

Pharmaceutical compositions comprising TITLE:

metformin and glibenclamide for the treatment

of type-II diabetes mellitus

INVENTOR (S): Tosetti, Alessandro; Guiducci, Mauro; Viti, Giovanni PATENT ASSIGNEE(S):

Menarini International Operations Luxembourg S.A.,

Luxembourg

SOURCE: PCT Int. Appl., 30 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

```
PATENT NO.
               KIND DATE
                                    APPLICATION NO. DATE
                     _____
                                     _____
                      20030605
WO 2003045355
                A1
                                     WO 2002-EP13497 20021129
   W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
       CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
       GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
       LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
       PL, PT, RO, RU, SC, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT,
       TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ,
       MD, RU, TJ, TM
   RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
       CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
       PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
       NE, SN, TD, TG
```

PRIORITY APPLN. INFO.:

IT 2001-FI230 A 20011129

Orally administrable pharmaceutical compns. in the form of tablets , comprising glibenclamide and metformin, or pharmaceutically acceptable salts thereof, as active ingredients, maintained sep. from one another within the same compn., are described for the treatment of type-II diabetes mellitus.

REFERENCE COUNT: THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 49 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2000:547376 HCAPLUS

DOCUMENT NUMBER:

133:155439

TITLE:

Controlled release tablets containing

biguanide and sulfonylurea

INVENTOR(S):

Chen, Chih-ming; Cheng, Xiu Xiu; Chou, Joseph; Jan,

Steve

PATENT ASSIGNEE(S):

Andrx Corporation, USA

SOURCE:

U.S., 9 pp. CODEN: USXXAM

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6099862	Α	20000808	US 1998-143876	19980831
CA 2341908	AA	20000309	CA 1999-2341908	19990831
JP 2003520759	T2	20030708	JP 2000-567214	19990831
US 6284275	B1	20010904	US 2000-590807	20000609
PRIORITY APPLN. INFO.:			US 1998-143876 A	19980831

WO 1999-US19978 W 19990831

A controlled release tablet contg. antihyperglycemic drug (that AB decreases hepatic glucose prodn.) and a hypoglycemic drug (that stimulates the release of insulin from the pancreas), that does not contain an expanding or gelling polymer layer, comprises a core of both the drugs, a semipermeable coating membrane surrounding the core and at least one passageway in the membrane to allow the drugs to be released from the core. A controlled release tablet contg. 500 mg metformin-HCl and 5 mg glipizide and having the following formulation was prepd.: metformin-HCl 87.77, glipizide 0.88, Povidone 6.31, sodium lauryl sulfate 4.54, and Mg stearate 0.50%. granules contg. the above formulation were compressed into tablets and coated with cellulose acetate 85, triacetin 5, and PEG 10%. 69 THERE ARE 69 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 50 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2003:334961 HCAPLUS

DOCUMENT NUMBER:

138:343914

TITLE:

Optimal polymer mixtures for gastric retentive

tablets

INVENTOR(S):

Gusler, Gloria; Berner, Bret; Chau, Mei; Padua, Aimee

Depomed, Inc., USA

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 22 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO.
                   KIND DATE
                                         APPLICATION NO. DATE
    <u>----</u>
                          _____
                                         _____
    WO 2003035177
                           20030501
                                         WO 2002-US33968 20021022
                     A2
                     A3
                           20030814
    WO 2003035177
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
            TJ, TM
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            CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
            PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
            NE, SN, TD, TG
    US 2003104053
                     A1
                          20030605
                                         US 2001-29134
                                                          20011025
                                      US 2001-29134
                                                     A 20011025
PRIORITY APPLN. INFO.:
```

Unit dosage form tablets for the delivery of pharmaceuticals are formed of the pharmaceutical dispersed in a solid unitary matrix that is formed of a combination of PEG and hydroxypropyl Me cellulose. The combination offers unique benefits in terms of release rate control and reproducibility while allowing both swelling of the tablet to effect gastric retention and gradual disintegration of the tablet to clear the tablet from the gastrointestinal tract after release of the drug has occurred. Thus, tablets contained gabapentin 60.0, PEG 24.3, HPMC 14.7, and Mg stearate 1.0%.

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L19 ANSWER 51 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
ACCESSION NUMBER: 2003:42092 HCAPLUS
DOCUMENT NUMBER:
                       138:112443
                       Tablet compositions for poorly-compressible
TITLE:
                       pharmaceuticals
INVENTOR(S):
                       Matharu, Amol Singh; Patel, Mahendra R.
                       Geneva Pharmaceuticals, Inc., USA
PATENT ASSIGNEE(S):
SOURCE:
                       PCT Int. Appl., 20 pp.
                       CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
                                        APPLICATION NO. DATE
                 KIND DATE
    PATENT NO.
    WO 2003004009 A1 20030116 WO 2002-US20323 20020627
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
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            TJ, TM
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            CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                    A1 20030130 US 2002-183881 20020627
    US 2003021841
                                     US 2001-302613P P 20010702
PRIORITY APPLN. INFO.:
    The present invention relates to a process for prepg. tablet
    dosage forms of poorly-compressible pharmaceuticals and to tablet
    dosage forms. The process is esp. useful for prepg. tablets of
    the poorly-compressible drug metformin-HCl. Thus,
     tablets contained metformin-HCl 500, HPMC 320, stearyl
    alc. 200, and Mg stearate mg/unit.
REFERENCE COUNT:
                              THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
                        5
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L19 ANSWER 52 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
ACCESSION NUMBER: 1999:613651 HCAPLUS
DOCUMENT NUMBER:
                       131:233581
TITLE:
                      Biphasic controlled-release delivery system for high
                       solubility pharmaceuticals
INVENTOR(S):
                       Timmins, Peter; Dennis, Andrew B.; Vyas, Kiren A.
PATENT ASSIGNEE(S):
                       Bristol-Myers Squibb Company, USA
                       PCT Int. Appl., 43 pp.
SOURCE:
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:
    PATENT NO. KIND DATE
                                        APPLICATION NO. DATE
    WO 9947128 A1 19990923 WO 1999-US5233 19990310
        W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
            DK, EE, ES, FI, GB, GE, GH, GM, HU, ID, IL, IS, JP, KE, KG, KP,
            KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO,
            NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA,
            UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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            ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
            CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
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CA 2320900

AA 19990923

CA 1999-2320900 19990310

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AU 1999-31828
                                                           19990310
                           19991011
    AU 9931828
                     A1
                    B2 20010809
    AU 736951
                                        EP 1999-913842 19990310
    EP 1063973
                     A1 20010103
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, FI
                                          BR 1999-8911
                                                         19990310
    BR 9908911
                           20011002
                                          JP 2000-536368 19990310
                      T2 20020305
    JP 2002506812
                                       US 1998-44446 A 19980319
PRIORITY APPLN. INFO.:
                                       WO 1999-US5233
                                                      W 19990310
    A biphasic controlled-release delivery system for pharmaceuticals which
AB
    have high water soly., such as the antidiabetic metformin-HCl,
    is provided which provides a dosage form that has prolonged gastric
    residence and includes (1) an inner solid particulate phase formed of
    substantially uniform granules contg. a pharmaceutical having a high water
     soly. and .gtoreq.1 hydrophilic polymer, .gtoreq.1 hydrophobic polymer,
    and/or .qtoreq.1 hydrophobic material such as waxes, fatty alcs., and/or
     fatty acid esters, and (2) an outer solid continuous phase in which the
    granules of the inner solid particulate phase are embedded and dispersed.
     The outer solid continuous phase includes .gtoreq.1 hydrophilic polymer,
     .gtoreq.1 hydrophobic polymer, and/or .gtoreq.1 hydrophobic material such
     as waxes, fatty alcs., and/or fatty acid esters, which may be compressed
     into tablets or filled into capsules. Methods for
     forming the biphasic controlled release delivery system and using it for
     treating diabetes are also provided. Thus, 500 g metformin-HCl
     was granulated with a dispersion of 25 g ethylcellulose in 100 mL 95%
     EtOH, dried, sieved, blended with hydroxypropylmethylcellulose 2208 USP
     351.5, hydroxypropylmethylcellulose 2910 USP 10, microcryst.
     cellulose 100.5 g, and 1% Mg stearate, and compressed into
     biphasic tablets each contg. 500 mg metformin-HCl.
     The percentage of metformin-HCl released from these
     tablets during in vitro testing was 38.1% after 1 h and 79.7%
     after 4 h.
                              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                        3
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L19 ANSWER 53 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
                      2002:122783 HCAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                        136:172785
                        Pharmaceutical composition comprising
TITLE:
                       metformin and a 5-phenoxyalkyl-2,4-
                        thiazolidinedione-type derivative
                        Moinet, Gerard; Botton, Gerard; Mesangeau, Didier
INVENTOR(S):
PATENT ASSIGNEE(S):
                        Merck Patent G.m.b.H., Germany
                        PCT Int. Appl., 18 pp.
SOURCE:
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
     PATENT NO. KIND DATE APPLICATION NO. DATE
     WO 2002011721 A1 20020214 WO 2001-EP8512 20010724
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
             RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
             UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
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BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

FR 2000-10362 20000804

FR 2812547

FR 2812547

A1

B1

20020208

20021031

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AU 2001-82010
    AU 2001082010
                     A5
                           20020218
                                                           20010724
    EP 1305025
                      A1
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                                         EP 2001-960539
                                                           20010724
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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                                          BR 2001-12915
    BR 2001012915
                     Α
                           20030708
                                                           20010724
    NO 2003000518
                      Α
                           20030203
                                          NO 2003-518
                                                           20030203
                                       FR 2000-10362
PRIORITY APPLN. INFO.:
                                                        A 20000804
                                       WO 2001-EP8512
                                                        W 20010724
OTHER SOURCE(S):
                        MARPAT 136:172785
    The present invention relates to an oral pharmaceutical compn. comprising,
     as active ingredients, metformin, optionally in the form of one
    of its pharmaceutically acceptable salts, and a 5-phenoxyalkyl-2,4-
    thiazolidinedione-type deriv. (I) for treatment of non-insulin-dependent
    diabetes. The wt. ratio of metformin or its salt to the compd.
    I, e.g., 5-[2-(4-cyanophenoxy)ethyl]thiazolidine-2,4-dione (CRE 16336),
    varies from 1:1 to 40:1. For example, a tablet was prepd.
    contg. metformin 850 mg, CRE 16336 50 mg, lactose 99 mg,
    hydroxypropyl cellulose 35 mg, sodium croscarmellose
     55 mg, and magnesium stearate 11 mg. The metformin and CRE
    16336 combination brings about normalization of the glycemia at doses
    where, given sep., these two products are without effect on the
    hyperglycemia.
REFERENCE COUNT:
                        3
                              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L19 ANSWER 54 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
                        2001:338335 HCAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                        134:344604
                        Antidiabetic formulation containing metformin
TITLE:
                        and sulfonylurea
INVENTOR (S):
                        Piper, Beth Anne
                        Bristol-Myers Squibb Company, USA
PATENT ASSIGNEE(S):
                        PCT Int. Appl., 76 pp.
SOURCE:
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                      APPLICATION NO. DATE
    PATENT NO.
                   KIND DATE
                    ____
                    A2
                                         WO 2000-US28467 20001013
    WO 2001032158
                           20010510
    WO 2001032158
                     A3 20020829
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
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            LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
             SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
            YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
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                                         US 1999-432465
    US 2002177602
                     A1
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    US 6586438
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    EP 1253944
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                                          EP 2000-970913
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    BR 2000015295
                           20030624
                                          BR 2000-15295
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    NO 2002002086
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                           20020624
                                          NO 2002-2086
                                                           20020502
                           20030228
                                          BG 2002-106732
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                                                           20020522
                           20030625
                                          LT 2002-62
    LT 5025
                      В
                                                           20020524
                                       US 1999-432465
PRIORITY APPLN. INFO.:
                                                        A 19991103
                                       WO 2000-US28467 W 20001013
    A low dose antidiabetic formulation adapted for treating e.g., Type II
AB
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diabetes contains a combination of metformin (at <800 mg/day) and at least 1 other antidiabetic agent such as a sulfonylurea. combination provides at least about substantially equiv. efficacy in treating diabetes as do antidiabetic formulations contg. metformin employed in dosages prescribed in generally accepted medical practice for first line therapy in treating diabetes, but with substantially reduced side effects, such as hypoglycemia and/or gastrointestinal distress. A method for treating diabetes in drug naive human patients is also provided employing the above formulation to reduce insulin resistance and/or post-prandial glucose excursion and/or Hb 1Ac, and/or increase post-prandial insulin, thereby treating the diabetes. tablets contained metformin-HCl 250.0, glyburide 1.25, croscarmellose sodium 7.00, Povidone 10.00, microcryst. cellulose 28.25, Mg stearate 2.25, and HPMC film-coating 6 mg. The effectiveness of this combination drug in producing hypoglycemia was demonstrated clin.

L19 ANSWER 55 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2002:845479 HCAPLUS

DOCUMENT NUMBER:

137:342124

TITLE:

Biphasic controlled-release delivery systems for high

solubility pharmaceuticals

INVENTOR(S):

Timmins, Peter; Dennis, Andrew B.; Vyas, Kiren A.

Bristol-Myers Squibb Company, USA

PATENT ASSIGNEE(S): SOURCE:

U.S., 16 pp., Cont.-in-part of U.S. Ser. No. 44,446,

APPLICATION NO. DATE

abandoned.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE _____ -----B1 20021105 US 1999-39810, US 1998-44446 B2 19980319 US 6475521 PRIORITY APPLN. INFO.: A biphasic controlled release delivery system for pharmaceuticals which have high water soly., such as the antidiabetic, metformin-HCl, provides a dosage form that has prolonged gastric residence so that a dosing regimen of at least one gram metformin, once daily, may be achieved while providing effective control of plasma glucose. delivery system includes (1) an inner solid particulate phase formed of substantially uniform granules contg. a pharmaceutical having a high water soly., and 1 or more hydrophilic polymers, 1 or more hydrophobic polymers and/or one or more hydrophobic materials such as 1 or more waxes, fatty alcs. and/or fatty acid esters, and (2) an outer solid continuous phase in which the above granules of inner solid particulate phase are embedded and dispersed throughout, the outer solid continuous phase including hydrophilic polymers, hydrophobic polymers and/or hydrophobic materials such as waxes, fatty alcs. and/or fatty acid esters, which may be compressed into tablets or filled into capsules. Methods for forming the so-described biphasic controlled release delivery system and using such biphasic controlled release delivery system for treating diabetes are also provided. Et cellulose N10 NF (25 g) was dissolved/dispersed in 100 mL ETOH. This dispersion was gradually added to 500 g metformin-HCl in a planetary mixer to produce a uniform damp granulation. The granulation was dried at 55.degree. for 1 h and passed through a 0.8-mm aperture screen to break down agglomerates. The metformin-Et cellulose granules (541 g) were blended with 351.5 g hydroxypropyl Me cellulose 2208 USP (100,000 cps grade), 10 g hydroxypropyl Me cellulose 2910 USP, and 100.5 g microcryst. cellulose in a planetary mixer for 10 min. Finally this mix was lubricated with 1% MG stearate and compressed into capsule-shaped tablets, each contg. 500 mg

metformin-HCl.

REFERENCE COUNT: 49 THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 56 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:152387 USPATFULL

TITLE: OPTIMAL POLYMER MIXTURES FOR GASTRIC RETENTIVE

TABLETS

INVENTOR(S): Gusler, Gloria, Cupertino, CA, UNITED STATES

> Berner, Bret, El Granada, CA, UNITED STATES Chau, Mei, Sunnyvale, CA, UNITED STATES Padua, Aimee, Daly City, CA, UNITED STATES

PATENT ASSIGNEE(S): DepoMed, Inc., Menlo Park, CA (U.S. corporation)

NUMBER KIND DATE -----

PATENT INFORMATION: US 2003104053 A1 20030605 APPLICATION INFO.: US 2001-29134 A1 20011025 (10)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: TOWNSEND AND TOWNSEND AND CREW, LLP, TWO EMBARCADERO

CENTER, EIGHTH FLOOR, SAN FRANCISCO, CA, 94111-3834

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 3 Drawing Page(s)

LINE COUNT: 705

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Unit dosage form tablets for the delivery of pharmaceuticals

are formed of the pharmaceutical dispersed in a solid unitary matrix

that is formed of a combination of poly(ethylene oxide) and

hydroxypropyl methylcellulose. The combination offers unique benefits in terms of release rate control and reproducibility while allowing both

swelling of the tablet to effect gastric retention and gradual

disintegration of the tablet to clear the tablet

from the gastrointestinal tract after release of the drug has occurred.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 57 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:81525 USPATFULL

TITLE:

Pharmaceutical composition comprising a combination of

metformin and fibrate, and its use for the preparation of medicines intended to reduce

hyperglycaemia

Bonhomme, Yves, Charbonnieres les Bains, FRANCE INVENTOR(S):

Briet, Philippe, Lyons, FRANCE

PATENT ASSIGNEE(S): Merck Patent Gesellschaft mit beschrankter Haftung,

Darnstadt, GERMANY, FEDERAL REPUBLIC OF (non-U.S.

corporation)

NUMBER KIND DATE -----US 6372790 B1 20020416 PATENT INFORMATION: WO 9940904 19990819 US 2000-601618 APPLICATION INFO.: 20001130 (9) WO 1999-EP614 19990130

20001130 PCT 371 date

NUMBER DATE

-----PRIORITY INFORMATION: FR 1998-1709 19980212

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Weddington, Kevin E. LEGAL REPRESENTATIVE: Millen, White, Zelano & Branigan, P.C.

NUMBER OF CLAIMS: 21 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT: 363

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A pharmaceutical composition comprising: (i) metformin, optionally in the form one of its pharmaceutically acceptable salts; (ii) a fibrate selected from fenofibrate and bezafibrate; and optionally one or more pharmaceutically acceptable excipients, is suitable for use

in the treatment of non-insulin-dependent diabetes.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 58 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:122779 HCAPLUS

DOCUMENT NUMBER: 136:172783

TITLE: Liquid formulation of metformin INVENTOR(S): Chandran, Ravi; Gogia, Ashish PATENT ASSIGNEE(S): Ranbaxy Signature LLC, USA

SOURCE: PCT Int. Appl., 42 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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KIND DATE
                                       APPLICATION NO. DATE
    PATENT NO.
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                                        ______
                                       WO 2001-IB1409 20010807
    WO 2002011716
                    A2
                          20020214
    WO 2002011716
                    A3
                          20020711
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
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            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
            RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
            UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
            BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                    A5
                                       AU 2001-76598
    AU 2001076598
                          20020218
                                                        20010807
    US 2002040063
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                          20020404
                                        US 2001-923491
                                                        20010807
    US 6559187
                     B2
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                                        BR 2001-13102
                                                         20010807
    BR 2001013102
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                          20030708
                                        US 2003-382442
                                                         20030306
    US 2003149111
                          20030807
                     A1
                                     US 2000-223391P P 20000807
PRIORITY APPLN. INFO.:
                                     US 2001-923491 A1 20010807
                                                    W 20010807
                                     WO 2001-IB1409
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An oral lig. compn. useful for treating hyperglycemia and diabetes AB comprises a therapeutically effective amt. of metformin or its pharmaceutically acceptable salt in a liq. carrier, i.e., water. compn. further comprises a sweetener that does not increase the blood glucose level of a subject after ingestion, alkyl hydroxyethyl cellulose, a polyhydroxy alc., and a mineral acid and a bicarbonate salt to maintain a pH of 4.0-9.0. The compn. addnl. comprises an antihyperglycemic agent, e.g., glyburide or glipizide. For example, to 60 L of purified water, heated to 40.degree., a mixt. of 1.9 kg of polyethylene glycol and 142.5 g hydroxyethyl cellulose (Natrosol 250 HX) was added. Then metformin-HCl (19 kg), followed by 1.188 kg calcium saccharin, 114 g citric acid, 211.28 g potassium benzoate, and addnl. polyethylene glycol (9.5 kg) were slowly added to the mixt., while maintaining the temp. of 40.degree.. A 70% soln. of sorbitol (in water) (76 kg) was pumped slowly to the tank maintained at 40.degree., and addnl. polyethylene glycol (21.85 kg) and cherry flavor (190 g) were

added to the tank and mixed. The contents of the tank were cooled to 30.degree., and addnl. water was added until the vol. was 190 L to obtain a metformin-HCl liq. formulation. The liq. formulation of the present invention contg. metformin or its pharmaceutically acceptable salt has several advantages over a solid formulation. Unlike the solid formulation, the liq. formulation can be administered to children and adults who have difficulty swallowing large size tablets. Thus, the liq. formulation facilitates patient compliance. Moreover, the liq. formulation showed to be safer and potentially exhibits less adverse effects than if the metformin or its salts were in a different formulation.

L19 ANSWER 59 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:152393 USPATFULL

Controlled release tablets of TITLE:

metformin

Chawla, Manish, Rohini, INDIA INVENTOR(S):

Raghuvanshi, Rajeev S., New Delhi, INDIA

Rampal, Ashok, Amritsar, INDIA

NUMBER KIND DATE _____ ____ US 2003104059 A1 20030605 US 2002-289070 A1 20021106 (10) PATENT INFORMATION:

APPLICATION INFO.:

NUMBER DATE _____

PRIORITY INFORMATION: IN 2001-11342001 20011106

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Ranbaxy Pharmaceuticals Inc., Suite 2100, 600 College

Road East, Princeton, NJ, 08540

NUMBER OF CLAIMS: 22 EXEMPLARY CLAIM: 1 LINE COUNT: 363

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Controlled-release metformin and processes for their

preparation, using a combination of non-ionic and anionic hydrophilic polymers, wherein the total hydrophilic polymer concentration is at least about 16% by weight of the composition.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 60 OF 256 USPATFULL on STN

2003:106816 USPATFULL ACCESSION NUMBER:

Combination of FBPase inhibitors and antidiabetic TITLE:

agents useful for the treatment of diabetes

INVENTOR(S): van Poelje, Paul D., La Jolla, CA, UNITED STATES

Erion, Mark D., Del Mar, CA, UNITED STATES

Fujiwara, Toshihiko, UNITED STATES

NUMBER KIND DATE -----US 2003073728 A1 20030417 US 2001-900364 A1 20010705 (9) PATENT INFORMATION: APPLICATION INFO.:

NUMBER DATE ______

US 2000-216531P 20000706 (60) PRIORITY INFORMATION: US 2000-215126P 20000629 (60)

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

BROBECK, PHLEGER & HARRISON LLP, 12390 EL CAMINO REAL, LEGAL REPRESENTATIVE:

SAN DIEGO, CA, 92130

NUMBER OF CLAIMS: 114 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 1 Drawing Page(s)

LINE COUNT: 12671

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A combination therapy of at least one FBPase inhibitor and at least one other antidiabetic agent is disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 61 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:928247 HCAPLUS

DOCUMENT NUMBER: 138:333

TITLE: Method for treating type 2 diabetes with low-dose

combination of metformin and glyburide

INVENTOR(S): Piper, Beth Anne

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 26 pp., Cont.-in-part of U.S.

Ser. No. 432,465. CODEN: USXXCO

DOCUMENT TYPE: Patent English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

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APPLICATION NO. DATE
    PATENT NO. KIND DATE
                                      _____
    -----
    US 2002183345 A1 20021205
                                      US 1999-460920 19991214
                   A1 20021128
                                      US 1999-432465 19991103
    US 2002177602
                   B2 20030701
    US 6586438
    WO 2001032157
                   A2 20010510
                                      WO 2000-US28311 20001013
                 A2 20020124
    WO 2001032157
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
           CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
           HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
           LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
           SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
           YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
           DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
           CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    EP 1229918
                   A2 20020814 EP 2000-972122 20001013
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
           IE, SI, LT, LV, FI, RO, MK, CY, AL
    JP 2003519621
                   T2 20030624
                                      JP 2001-534362 20001013
                         20030715
                                      BR 2000-15294 20001013
    BR 2000015294
                    Α
    NO 2002002087
                    Α
                       20020624
                                      NO 2002-2087
                                                     20020502
    BG 106733
                    A 20030228
                                     BG 2002-106733 20020522
                                      LT 2002-62
                    B 20030625
                                                     20020524
    LT 5025
                                    US 1999-432465 A2 19991103
PRIORITY APPLN. INFO.:
                                    US 1999-460920 A 19991214
                                    WO 2000-US28311 W 20001013
```

AB A method is provided for first line treatment of type 2 diabetes employing a combination of metformin and glyburide. A method for treating diabetes in drug naive human patients is also provided employing the above formulation to reduce insulin resistance and/or post-prandial glucose excursion and/or Hb Alc, and/or increase post-prandial insulin, thereby treating the diabetes. Hydroxypropylmethylcellulose film-coated tablets of metformin HCl and glyburide were prepd. and tested in drug naive patients with type 2 diabetes mellitus who have had inadequate glycemic control with diet and exercise. A low dose metformin-glyburide (250 mg/1.25 mg) formulation achieved glycemic control at least essentially equiv. to a high dose metformin -glyburide (500 mg/2.5 mg) formulation but with reduced incidence of side

effects.

PATENT INFORMATION:

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L19 ANSWER 62 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
                         1999:613646 HCAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         131:233580
TITLE:
                         Controlled release oral tablet having a
                         unitary core
                         Cheng, Xiu Xiu; Chen, Chih-Ming; Jan, Steve; Chou,
INVENTOR(S):
                         Joseph
                         Andrx Pharmaceuticals, Inc., USA
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 30 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
                         English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                    KIND DATE
                                         APPLICATION NO. DATE
                    ----
                      A1
                            19990923
                                          WO 1999-US6024 19990319
     WO 9947125
        W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG,
             KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
             UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
             ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                         US 1998-45330
                                                            19980320
                            20000808
     US 6099859
                       Α
                                           CA 1999-2324493 19990319
     CA 2324493
                            19990923
                       AA
     AU 9931019
                            19991011
                                           AU 1999-31019
                                                            19990319
                       A1
     AU 739226
                       B2
                            20011004
     EP 1063971
                                           EP 1999-912705
                                                            19990319
                      A1
                            20010103
         R: CH, DE, DK, ES, FR, GB, IT, LI, NL, SE
                            20020305
                                          JP 2000-536365
                                                            19990319
     JP 2002506810
                     T2
                                           US 2000-726193
                                                            20001129
     US 2001024659
                            20010927
                       A1
     US 2002064556
                            20020530
                                           US 2001-16556
                                                            20011101
                       A1
     US 6495162
                       B2
                            20021217
                                        US 1998-45330
                                                         A 19980320
PRIORITY APPLN. INFO.:
                                        WO 1999-US6024
                                                         W 19990319
                                        US 2000-594637
                                                         A1 20000615
     A controlled release antihyperglycemic tablet that does not
AB
     contain an expanding polymer comprises a core contg. the antihyperglycemic
     drug, a semipermeable membrane coating the core and at least one
     passageway in the membrane. A core was prepd. contg. metformin
     -HCL 90.54, Povidone 4.38, Na3PO4 4.58, and Mg stearate 0.5% and a
     sustained release coting comprised cellulose acetate 85,
     triacetin 5, and PEG 400 10%.
                               THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                         1
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L19 ANSWER 63 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
                         2003:490987 HCAPLUS
ACCESSION NUMBER:
                         139:57931
DOCUMENT NUMBER:
                         Antidiabetic formulation containing metformin
TITLE:
                         and glipizide
                         Li, Danping; Phusanti, Lawan; Desai, Divyakant S.
INVENTOR(S):
                         Bristol-Myers Squibb Company, USA
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 28 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
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APPLICATION NO. DATE
                   KIND DATE
    PATENT NO.
                                          -----
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                                          WO 2002-US39140 20021209
    WO 2003051293
                    A2
                           20030626
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD,
            RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
            CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
            PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
            MR, NE, SN, TD, TG
                     A1 20030724
                                          US 2001-23533
                                                          20011217
    US 2003139461
                                       US 2001-23533
                                                       A 20011217
PRIORITY APPLN. INFO.:
    An antidiabetic pharmaceutical formulation is provided, esp. adapted for
    treating Type II diabetes, which includes a combination of
    metformin and glipizide in a manner to control moisture in the
    formulation so that the glipizide does not hydrolyze, yet the
    metformin is compressible, if necessary. Excipients that are used
     in the formulations are microcryst. cellulose, Povidone,
     Croscarmellose sodium, Mg stearate, and HPMC.
L19 ANSWER 64 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
                        2002:793392 HCAPLUS
ACCESSION NUMBER:
                        137:299938
DOCUMENT NUMBER:
                        Timed pulse release composition containing swellable
TITLE:
                        core and polymeric coat
                        Shanghvi, Dilip Shantilal; Dharmadhikari, Nitin
INVENTOR(S):
                        Bhalachandra; Zala, Yashoraj Rupsinh; Khanna, Satish
                        Sun Pharmaceutical Industries Limited, India
PATENT ASSIGNEE(S):
                        PCT Int. Appl., 21 pp.
SOURCE:
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                     KIND DATE
                                          APPLICATION NO. DATE
                           -----
                     ____
                      A2
                           20021017
                                          WO 2002-IN107
                                                           20020409
     WO 2002080887
                           20021128
                      Α3
     WO 2002080887
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
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CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
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        LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
        PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
        UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
        TJ, TM
    RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
        CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
        BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                     WO 2002-IN203
                                                      20021008
                     20030417
WO 2003030920
                 A1
                       20030626
WO 2003030920
                 C2
       AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
        CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
        GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
        LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
        PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
        UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD,
        RU, TJ, TM
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
    CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
    PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
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NE, SN, TD, TG

PRIORITY APPLN. INFO.: IN 2001-MU325 A 20010410

IN 2001-MU984 A 20011008 WO 2002-IN107 A 20020409

The present invention provides a timed pulse release compn. comprising: AΒ (a) a core compn. comprising a therapeutically active agent, a swelling agent, and optionally water sol. compd.(s) for inducing osmosis, and (b) a coat compn. comprising one or more film forming polymers. Upon imbibing fluid from the surrounding, the core swells, and the coat ruptures to release in a pulse the therapeutically active agent in a reliable manner at about a predetd. time; the reliable manner of rupture comprises rupturing of 36 tablets out of a total of 36 tablets at about the predetd. time when tested by subjecting the tablets to USP dissoln. test using an aq. media at 37.degree., in a USP Type I or Type II app. at about 50-100 rpm. For example, a timed pulse release tablet was prepd. contg. (as core) metformin hydrochloride 500.0 mg, AcDiSol 50.0 mg, corn starch (10% starch paste) 17.0 mg, microcryst. cellulose 13.5 mg, colloidal silica 13.5 mg, and magnesium stearate 6.0 mg, and (as a coat) Et cellulose 40.7 mg, and hydroxypropyl Me cellulose 16.3 mg. Tablets released the metformin as a pulse after the rupture of the coat at a predetd. time (about 1-1.3 h).

L19 ANSWER 65 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:284364 HCAPLUS

DOCUMENT NUMBER:

138:44601

TITLE:

Functionality testing of a multifunctional directly

compressible adjuvant containing lactose, polyvinylpyrrolidone, and croscarmellose

sodium

AUTHOR (S):

Gohel, Mukesh C.; Jogani, Pranav D.

CORPORATE SOURCE:

Lallubhai Motilal College of Pharmacy, Ahmedabad, 380

009, India

SOURCE:

Pharmaceutical Technology North America (2002), 26(3),

64,66,68,70,72,74,76,78,80,82 CODEN: PTNABQ; ISSN: 1534-2131

PUBLISHER:

Advanstar Communications, Inc.

DOCUMENT TYPE:

Journal LANGUAGE: English

A directly compressible multifunctional adjuvant contg. lactose, polyvinylpyrrolidone, and croscarmellose sodium was prepd. by using a simple solvent-free method. The flowability and compressibility of the agglomerates obtained were significantly superior to those of lactose monohydrate. The agglomerates exhibited good diln. potential and were sensitive to high humidity. Tablets prepd. by using herbal drugs (Glycyrrhiza and turmeric) and synthetic drugs such as metformin-HCl and acetaminophen were satisfactory.

L19 ANSWER 66 OF 256 USPATFULL on STN

ACCESSION NUMBER:

1999:78774 USPATFULL

TITLE:

Glibenclamide-metformin combination for the treatment of diabetes mellitus of type II

INVENTOR(S):

Barelli, Giulio, Pisa, Italy De Regis, Massimo, Pisa, Italy

PATENT ASSIGNEE(S):

Abiogen Pharma s.r.l., Italy (non-U.S. corporation)

	NUMBER	KIND	DATE	
PATENT INFORMATION: U	JS 5922769		19990713	
V	<i>1</i> 0 9717975		19970522	
APPLICATION INFO.: U	JS 1998-29371		19980513	(9)

WO 1996-EP4860 19961107

> 19980513 PCT 371 date 19980513 PCT 102(e) date

NUMBER DATE ------

PRIORITY INFORMATION: IT 1995-MI2337 19951114

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Henley, III, Raymond LEGAL REPRESENTATIVE: Nixon & Vanderhye

NUMBER OF CLAIMS: 1 EXEMPLARY CLAIM: 509 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Non-insulin dependent diabetes mellitus in cases of secondary failure is

treated with a combination of glibenclamide and metformin.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 67 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:180234 USPATFULL

Pharmaceutical safety dosage forms TITLE:

Roberts, Richard H., Lakewood, NJ, UNITED STATES INVENTOR(S):

KIND DATE NUMBER ______

US 2003124061 A1 20030703 US 2003-339977 A1 20030110 (10)

PATENT INFORMATION:
APPLICATION INFO.:
DOCUMENT TYPE:
FILE SEGMENT: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: WOODCOCK WASHBURN LLP, ONE LIBERTY PLACE, 46TH FLOOR,

1650 MARKET STREET, PHILADELPHIA, PA, 19103

NUMBER OF CLAIMS: 228
EXEMPLARY CLAIM: 1
NUMBER OF DRAWINGS: 1 Drawing Page(s)
1140
1200 THIS PATENT

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Pharmaceutical safety dosage forms are provided which include a pharmaceutical and an antagonist to the pharmaceutical. The safety dosage forms are such that the antagonist has no significant bioavailability when the pharmaceutical safety dosage form is administered as intended. However, the antagonist is released and becomes bioavailable if the dosage form is disrupted. Methods of administering pharmaceuticals by providing pharmaceutical safety dosage

forms are also provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 68 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2001:114962 HCAPLUS

134:152671 DOCUMENT NUMBER:

Floating pharmaceutical composition comprising an TITLE:

active phase and a non-active phase

Besse, Jerome INVENTOR(S):

PATENT ASSIGNEE(S): Galenix Developpement, Fr. SOURCE: PCT Int. Appl., 33 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent French LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

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WO 2001010417 A1 20010215 WO 2000-FR2223 20000802
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
            HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
            LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
            SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
            YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
            CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                    FR 1999-10285
                    A1 20010209
                     B1 20011026
    FR 2797185
                     A1 20020522 EP 2000-956599 20000802
    EP 1206247
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL
    BR 2000013106 A 20020723 BR 2000-13106 20000802
NO 2002000572 A 20020403 NO 2002-572 20020205
PRIORITY APPLN. INFO.:
                                       FR 1999-10285 A 19990806
                                       WO 2000-FR2223 W 20000802
    The invention concerns a floating pharmaceutical compn. consisting of at
AΒ
    least a first phase comprising at least a high dose active principle
    combined with one or several carriers and at least a second phase
    comprising at least a gas-generating system. The invention also concerns
    tablets comprising such a pharmaceutical compn. and a method for
    prepg. such tablets. A programmed-release tablet
    contained metformin hydrochloride 51.33, Carbopol-974 3.02,
    hydroxypropyl cellulose 4.53, magnesium stearate 0.06% in the
    active layer; and hydroxypropylmethyl cellulose 24.64,
    monosodium citrate 7.23 in the non-active layer.
                              THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                        5
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L19 ANSWER 69 OF 256 USPATFULL on STN
                       2003:201447 USPATFULL
ACCESSION NUMBER:
                       Combinations comprising dipeptidylpeptidase-iv
TITLE:
                       inhibitor
                       Balkan, Bork, Madison, CT, UNITED STATES
INVENTOR(S):
                       Hughes, Thomas Edward, Somerville, NJ, UNITED STATES
                       Holmes, David Grenville, Binningen, SWITZERLAND
                       Villhauer, Edwin Bernard, Morristown, NJ, UNITED STATES
                                        KIND DATE
                            NUMBER
                       _____
                       US 2003139434 A1 20030724 US 2002-181169 A1 20021010 (10) WO 2001-EP590 20010119
PATENT INFORMATION:
APPLICATION INFO.:
                             NUMBER
                                          DATE
                       ______
                       US 2000-9489234 20000121
PRIORITY INFORMATION:
                       US 2000-9619262 20000719
DOCUMENT TYPE:
                       Utility
FILE SEGMENT:
                       APPLICATION
                       THOMAS HOXIE, NOVARTIS, PATENT AND TRADEMARK
LEGAL REPRESENTATIVE:
                       DEPARTMENT, ONE HEALTH PLAZA 430/2, EAST HANOVER, NJ,
                       07936-1080
NUMBER OF CLAIMS:
                       16
EXEMPLARY CLAIM:
LINE COUNT:
                       1581
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The invention relates to a combination which comprises a DPP-IV
       inhibitor and at least one further antidiabetic compound, preferably
```

selected from the group consisting of insulin signalling pathway

modulators, like inhibitors of protein tyrosine phosphatases (PTPases), non-small molecule mimetic compounds and inhibitors of glutamine-fructose-6-phosphate amidotransferase (GFAT), compounds influencing a dysregulated hepatic glucose production, like inhibitors of glucose-6-phosphatase (G6Pase), inhibitors of fructose-1,6bisphosphatase (F-1,6-BPase), inhibitors of glycogen phosphorylase (GP), glucagon receptor antagonists and inhibitors of phosphoenolpyruvate carboxykinase (PEPCK), pyruvate dehydrogenase kinase (PDHK) inhibitors, insulin sensitivity enhancers, insulin secretion enhancers, .alpha.-glucosidase inhibitors, inhibitors of gastric emptying, insulin, and .alpha..sub.2-adrenergic antagonists, for simultaneous, separate or sequential use in the prevention, delay of progression or treatment of conditions mediated by dipeptidylpeptidase-IV (DPP-IV), in particular diabetes, more especially type 2 diabetes mellitus, conditions of impaired glucose tolerance (IGT), conditions of impaired fasting plasma qlucose, metabolic acidosis, ketosis, arthritis, obesity and osteoporosis; and the use of such combination for the cosmetic treatment of a mammal in order to effect a cosmetically beneficial loss of body weight.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 70 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2000:716155 HCAPLUS

DOCUMENT NUMBER: 134:285541

TITLE: New excipients in fast-release tablet

formulations

AUTHOR(S): Fang, Xiao-Ling; Yang, Min; Mu, Ni-La; Wang,

Xue-Liang; Zhang, Jin

CORPORATE SOURCE: Dept. of Pharmaceutics, Shanghai Medical University,

Shanghai, 200032, Peop. Rep. China

SOURCE: Zhongguo Yiyao Gongye Zazhi (2000), 31(6), 257-259

CODEN: ZYGZEA; ISSN: 1001-8255

PUBLISHER: Zhongguo Yiyao Gongye Zazhi Bianjibu

DOCUMENT TYPE: Journal LANGUAGE: Chinese

The water-sol. but poor-compressible drug metformin-HCl and the poor water-sol. but good-compressible drug ofloxacin were chosen as model drugs for test. The formulations were designed and evaluated using super-disintegrants (crosslinking PVP, crosslinking CMC-Na, L-HPC and CMS-Na), new binder PVP, and new filler microcryst. cellulose. The quality criteria such as granular property, compressibility, disintegration time and dissoln. for various formulations indicated that these new excipients could be used satisfactorily in fast-release tablets formulation.

L19 ANSWER 71 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:40435 USPATFULL

TITLE: Oral formulation comprising biguanide and an organic

acid

INVENTOR(S): Nishii, Hiroyuki, Osaka, JAPAN

Kobayashi, Hirohisa, Osaka, JAPAN Otoda, Kazuya, Takarazuka, JAPAN

PATENT ASSIGNEE(S): Sumitomo Pharmaceuticals Company, Limited, Osaka, JAPAN

(non-U.S. corporation)

NUMBER DATE

PRIORITY INFORMATION: JP 1998-136126 19980429

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Spear, James M.

LEGAL REPRESENTATIVE: Birch, Stewart, Kolasch & Birch, LLP

NUMBER OF CLAIMS: 12 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 0 Drawing Figure(s); 0 Drawing Page(s)

LINE COUNT: 302

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB An oral formulation comprising a biguanide and an organic acid has less

unpleasant tastes such as bitterness and saltiness.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 72 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2001:193965 USPATFULL

TITLE: Core formulation

INVENTOR(S): Adjei, Akwete L., Bridgewater, NJ, United States

Zhu, Yaping, Highland Park, International Patent

Institute

Cutie, Anthony J., Bridgewater, International Patent

Institute

NUMBER DATE

PRIORITY INFORMATION: US 2000-201233P 20000501 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Jerome Rosenstock, Frommer Lawrence & Haug LLP, 745

Fifth Avenue, New York, NY, 10151

NUMBER OF CLAIMS: 18
EXEMPLARY CLAIM: 1
LINE COUNT: 532

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention relates to a controlled-release combination drug product comprising troglitazone, e.g. its hydrochloride, and a biguamide, e.g.

metformin. In particular, the product comprises a core of metformin, at least a portion thereof has a layer or coat

thereon of troglitazone.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 73 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2003:609873 HCAPLUS

DOCUMENT NUMBER: 139:154910

TITLE: Manufacture of oral dosage forms delivering both

immediate-release and sustained-release drugs

INVENTOR(S): Lim, Jong C.; Shell, John N.

PATENT ASSIGNEE(S): Depomed, Inc., USA

SOURCE: U.S. Pat. Appl. Publ., 5 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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APPLICATION NO. DATE
PATENT NO.
               KIND DATE
                                US 2002-66146 20020201
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US 2003147952 A1 20030807
                                                  20020201
WO 2003066028
               A1 20030814
                                  WO 2003-US2809 20030128
      AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
       CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
       GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
       LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
       PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
       UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
       TJ, TM
   RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
       CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC,
       NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW,
       ML, MR, NE, SN, TD, TG
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PRIORITY APPLN. INFO.: US 2002-66146 A 20020201

A method is disclosed for manufg. a pharmaceutical tablet for oral administration, the tablet combining both immediate-release and prolonged-release modes of drug delivery and using an immediate-release drug that is either insol. in water or only sparingly sol. and is present in a very small amt. compared to the prolonged-release drug. The method involves the use of particles of the immediate-release drug that are equal to or less than 10 .mu. in diam., applied as a layer or coating over a core of the prolonged-release drug, the layer or coating being either the drug particles themselves, applied as an aq. suspension, or a solid mixt. contg. the drug in admixt. with a material that disintegrates rapidly in gastric fluid. The result in both cases is a high degree of uniformity in the proportions of the immediate-release and prolonged-release drugs, uniformity that is otherwise difficult to achieve in view of the insoly. of the immediate-release drug and its relatively small amt. compared to the prolonged-released drug. Tablets contg. metformin-HCl and glimepiride were prepd. contg. HPMC and PEG, using Polysorbate 80 solns.

L19 ANSWER 74 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1987:605192 HCAPLUS

DOCUMENT NUMBER: 107:205192

TITLE:

N, N-dimethylbiguanide p-chlorophenoxyacetate pharmaceutical preparation for treatment of

neuropathies

INVENTOR(S):

Hugelin, Andre; Thal, Claude

PATENT ASSIGNEE(S):

Fr.

SOURCE:

Fr. Demande, 8 pp.

CODEN: FRXXBL

DOCUMENT TYPE:

Patent French

LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PAT	TENT NO.	KIND	DATE	APPLICATION NO.	DATE
FR	2585572	A1	19870206	FR 1985-11664	19850731
FR	2585572	B1	19871231		
ΑU	8660768	A1	19870205	AU 1986-60768	19860731
ΑU	587054	B2	19890803		
ΕP	214017	A2	19870311	EP 1986-401717	19860731
ΕP	214017	A3	19890726		
EP	214017	B1	19920722		
	R: AT, B	E, CH, DE	, FR, GB,	IT, LI, LU, NL, SE	
JΡ	62116510	A2	19870528	JP 1986-181195	19860731
JP	07025671	B4	19950322		
ZA	8605735	A	19871125	ZA 1986-5735	19860731
AT	78397	E	19920815	AT 1986-401717	19860731

US 4835184 A 19890530 US 1986-893025 19860801 PRIORITY APPLN. INFO.: FR 1985-11664 19850731 EP 1986-401717 19860731

AB Neurotrophic pharmaceuticals contain a neurol. active quantity of N,N-dimethylbiguanide p-chlorophenoxyacetate (I). Effervescent tablets contained I 1500, corn starch 24, wheat starch 36, lactose 375, NaHCO3 12.6, tartaric acid 11.25, hydroxypropyl cellulose 18, Povidone C15 12, and a sugar glaze 120 mg. I was as effective as Isaxonine in regeneration of nerve fibers in rats, whereas N,N-dimethylbiguanide-HCl was ineffective.

L19 ANSWER 75 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2001:188740 USPATFULL

TITLE: Core formulation

INVENTOR(S): Adjei, Akwete L., Bridgewater, NJ, United States

Zhu, Yaping, Highland Park, NJ, United States Cutie, Anthony J., Bridgewater, NJ, United States

NUMBER DATE

PRIORITY INFORMATION: US 2000-201233P 20000501 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Jerome Rosenstock, Esq, c/o FROMMER LAWRENCE & HAUG

LLP, 745 Fifth Avenue, New York, NY, 10151

NUMBER OF CLAIMS: 17 EXEMPLARY CLAIM: 1

LINE COUNT: 544

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention relates to a controlled release combination drug product comprising troglitazone, e.g. its hydrochloride, and a biguamide, e.g.

metformin. In particular, the product comprises a core of

metformin. In particular, the product comprises a core of metformin, at least a portion thereof has a layer or coat

thereon of troglitazone.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 76 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:136590 USPATFULL

TITLE: Core formulation

INVENTOR(S): Adjei, Akwete L., Bridgewater, NJ, United States

Zhu, Yaping, Highland Park, NJ, United States Cutie, Anthony J., Bridgewater, NJ, United States

PATENT ASSIGNEE(S): Aeropharm Technology Incorporated, Edison, NJ, United

States (U.S. corporation)

NUMBER DATE

PRIORITY INFORMATION: US 2000-201057P 20000501 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: GRANTED

PRIMARY EXAMINER: Henley, III, Raymond

LEGAL REPRESENTATIVE: Frommer Lawrence & Haug LLP

NUMBER OF CLAIMS: 20

EXEMPLARY CLAIM: 1

0 Drawing Figure(s); 0 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 493

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention relates to a modulating formulation comprising pioglitazone hydrochloride and a biguamide, e.g. metformin. In particular, the product comprises a core of the biguamide, e.g.

metformin, at least a portion thereof has a layer or coat

thereon of pioglitazone.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 77 OF 256 USPATFULL on STN

ACCESSION NUMBER:

2001:218028 USPATFULL

TITLE:

Core formulation

INVENTOR(S):

Adjei, Akwete L., Bridgewater, NY, United States Zhu, Yaping, Highland Park, NJ, United States Cutie, Anthony J., Bridgewater, NJ, United States

NUMBER KIND DATE US 2001046515 A1 20011129 US 6524621 B2 20030225 PATENT INFORMATION: US 2001-784713 A1 20010215 (9) APPLICATION INFO.:

> NUMBER DATE

PRIORITY INFORMATION:

US 2000-201057P 20000501 (60)

DOCUMENT TYPE:

Utility APPLICATION

FILE SEGMENT:

LEGAL REPRESENTATIVE: JEROME ROSENSTOCK, FROMMER LAWRENCE & HAUG LLP, 745

Fifth Avenue, New York, NY, 10151

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

1

LINE COUNT:

493

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention relates to a modulating formulation comprising pioglitazone hydrochloride and a biguamide, e.g. metformin. In particular, the product comprises a core of the biguamide, e.g.

metformin, at least a portion thereof has a layer or coat

thereon of pioglitazone. PATENT

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 78 OF 256 USPATFULL on STN

ACCESSION NUMBER:

2001:29618 USPATFULL

TITLE:

Use of metformin to counteract weight gain

associated with valproate and other psychotropic

medications

INVENTOR(S):

Cottingham, Elizabeth Marie, 300 Warren Ave.,

Cincinnati, OH, United States 45219

Morrison, John Ainslie, 3740 Clifton Ave., Cincinnati,

OH, United States 45220

NUMBER KIND DATE ______ US 6194466 B1 20010227 US 1999-416330 19991012 PATENT INFORMATION:

APPLICATION INFO.:

19991012 (9)

NUMBER DATE -----

PRIORITY INFORMATION:

US 1998-104394P 19981015 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

Granted

PRIMARY EXAMINER:

Criares, Theodore J.

ASSISTANT EXAMINER:

Kim, J.

LEGAL REPRESENTATIVE:

Frost Brown Todd LLC

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

LINE COUNT:

224

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for minimizing the weight gain side effect associated with

Valproate treatment is disclosed. In this method, Metformin, a

biquanide compound, is concurrently administered to a patient taking the

Valproate therapy. A pharmaceutical composition containing the

combination of Valproate and Metformin is also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 79 OF 256 USPATFULL on STN

ACCESSION NUMBER:

2001:167766 USPATFULL

TITIE:

Core formulation comprising troglitazone and abiquanide

INVENTOR (S):

Cutie, Anthony J., Bridgewater, NJ, United States

PATENT ASSIGNEE(S):

Adjei, Akwete L., Bridgewater, NJ, United States

Aeropharm Technology Incorporated, Edison, NJ, United

States (U.S. corporation)

NUMBER -----

KIND DATE

PATENT INFORMATION:

US 6296874 B1 20011002 US 2000-703023 20001031 APPLICATION INFO.:

20001031 (9)

NUMBER DATE

PRIORITY INFORMATION:

______ US 2000-201233P 20000501 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

GRANTED

PRIMARY EXAMINER: GRANTED

PRIMARY EXAMINER: Spear, James M.

NUMBER OF CLAIMS:

LEGAL REPRESENTATIVE: Frommer Lawrence & Haug LLP

EXEMPLARY CLAIM:

17

LINE COUNT:

384

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention relates to a combination drug product comprising troqlitazone, e.g. its hydrochloride, and a biguamide, e.g. metformin. In particular, the product comprises a core of metformin, at least a portion thereof has a layer or coat thereon of troglitazone.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 80 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:391499 HCAPLUS

DOCUMENT NUMBER:

136:406855

TITLE:

SOURCE:

Medicine based on antihyperglycemic microcapsules with

prolonged release and method for preparing same Castan, Catherine; Meyrueix, Remi; Soula, Gerard

INVENTOR(S): PATENT ASSIGNEE(S):

Flamel Technologies, Fr. PCT Int. Appl., 28 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.

KIND DATE

APPLICATION NO. DATE

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WO 2001-FR3625
                                                          20011119
                    A2
                           20020523
    WO 2002039984
                    A3
                          20020711
    WO 2002039984
           AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA,
            UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
            CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                         20020524 FR 2000-14876 20001117
                    A1
    FR 2816840
                          20020527 AU 2002-20796 20011119
20030813 EP 2001-996365 20011119
    AU 2002020796
                      Α5
    EP 1333816
                     A2
           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                                                      A 20001117
                                      FR 2000-14876
PRIORITY APPLN. INFO.:
                                      WO 2001-FR3625 W 20011119
    The invention concerns an oral galenic form for prolonged release of
AB
    anti-hyperglycemic (metformin) active principles. Said medicine
    enables to obtain an efficient therapeutic protection over 24 h by
    overcoming the problems of bypass of the absorption window and the massive
    localized release of active principles. Therefor, said medicine comprises
    several thousand anti- hyperglycemic (metformin) microcapsules
    each consisting of a core comprising at least an anti- hyperglycemic agent
    and of a coating film applied on the core and enabling the
    prolonged release in vivo of the anti- hyperglycemic agent. Said
    microcapsules have a grain size distribution ranging between 50 and 100
     .mu.. The reproducibility of the transit kinetics and hence of
    bioavailability are very high. There results for the patient a lesser
    risk of hyperglycemic or hypoglycemic. The invention also concerns the
    prepn. of said medicine and the use of a plurality of said microcapsules
    for making an anti- hyperglycemic medicine. The invention is applicable
    to the treatment of type II diabetes. A soln. of 159.5 g stearic acid and
    159.5 g Et cellulose in 2870 g isopropanol at 50.degree. was
    sprayed on 700 g of metformin hydrochloride crystals (av. diam.
    100-200 .mu.m). The dissoln. rate of the granules thus obtained was 97.1%
    after 20 min.
L19 ANSWER 81 OF 256 USPATFULL on STN
ACCESSION NUMBER:
                       2001:152504 USPATFULL
                       Pharmaceutical compositions of vanadium biguanide
TITLE:
                       complexes and their use
INVENTOR(S):
                       Orvig, Chris, Vancover, Canada
                       McNeill, John H., Vancouver, Canada
                       The University of British Columbia, Vancouver, Canada
PATENT ASSIGNEE(S):
                       (non-U.S. corporation)
                           NUMBER
                                       KIND DATE
                       _____
                       US 6287586 B1 20010911
PATENT INFORMATION:
                       US 1999-396982
                                              19990915 (9)
APPLICATION INFO.:
                            NUMBER
                                          DATE
                       ______
PRIORITY INFORMATION:
                       US 1998-101074P 19980918 (60)
DOCUMENT TYPE:
                       Utility
FILE SEGMENT:
                       GRANTED
                       Webman, Edward J.
PRIMARY EXAMINER:
ASSISTANT EXAMINER:
                       Nguyen, Helen
                       Sherwood, Pamela J.Bozicevic, Field & Francis LLP
LEGAL REPRESENTATIVE:
NUMBER OF CLAIMS:
                       10
EXEMPLARY CLAIM:
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1 Drawing Figure(s); 1 Drawing Page(s)

NUMBER OF DRAWINGS:

LINE COUNT: 798

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Pharmaceutical compositions of vanadium biguanide complexes, and methods of use, are provided for the treatment of hyperglycemia and related disorders, e.g. hypertension, obesity, and lipid disturbances. The pharmaceutically active complexes of the invention comprise a biguanide chelant, preferably a 1-substituted biguanide chelant, capable of chelating vanadium to form a six-membered unsaturated vanadium-containing ring. The vanadium of the complex is coordinated with oxygen, sulphur or nitrogen, particularly oxygen coordinated. The complexes are formulated with a physiologically acceptable carrier. In a preferred embodiment, the complexes are formulated for oral administration.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 82 OF 256 USPATFULL on STN

ACCESSION NUMBER: 75:21254 USPATFULL

TITLE: Antihyperglycemic methods and compositions

INVENTOR(S): Kabbe, Hans-Joachim, Leverkusen, Germany, Federal

Republic of

Horstmann, Harald, Wuppertal-Elberfeld, Germany,

Federal Republic of

Plumpe, Hans, Wuppertal-Elberfeld, Germany, Federal

Republic of

Puls, Walter, Wuppertal-Elberfeld, Germany, Federal

Republic of

Petersen, Siegfried, Leverkusen, Germany, Federal

Republic of

PATENT ASSIGNEE(S): Bayer Aktiengesellschaft, Leverkusen, Germany, Federal

Republic of (non-U.S. corporation)

NUMBER KIND DATE
----US 3879541 19750422
US 1973-324218 19730116 (5)

APPLICATION INFO.: US 1973-324218 19730116 (5)
RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1971-118958, filed

on 25 Feb 1971, now abandoned And Ser. No. US

on 25 Feb 1971, now abandoned And Ser. No. US 1971-120332, filed on 2 Mar 1971, now abandoned

NUMBER DATE
----DE 1970-2009738 19700303

PRIORITY INFORMATION: DE 1970-2009738 19700303 DE 1970-2009743 19700303

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Waddell, Frederick E.

NUMBER OF CLAIMS: 14
EXEMPLARY CLAIM: 1
LINE COUNT: 470

PATENT INFORMATION:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The blood sugar level of hyperglycemic animals can be reduced through administration of an N.sup.1 -phenylbiguanide which is substituted in the N.sup.5 -position by the group CH.sub.2 R.sup.2 in which R.sup.2 is hydrogen, alkyl of 1 to 7 carbon atoms, alkoxyalkyl of 2 to 5 carbon atoms, cyclohexyl or vinyl, and optionally substituted by one or two groups in the phenyl ring. Solid, orally administered pharmaceutical compositions are also described. A typical embodiment is the use of N.sup.1 - (4-chlorophenyl)-N.sup.5 - (n-propyl) biguanide hydrochloride which can be administered in a tablet, capsule or dragee.

L19 ANSWER 83 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:237424 USPATFULL

Compositions for treating diabetes mellitus, methods of TITLE:

use and manufacturing process of the same

INVENTOR(S): Wang, Peng, Burlingame, CA, UNITED STATES

Lei, Lin, Melshan, CHINA

NUMBER KIND DATE ______

PATENT INFORMATION: US 2003165581 A1 20030904 US 2002-91371 A1 20020304 (10) APPLICATION INFO.:

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: WILSON SONSINI GOODRICH & ROSATI, 650 PAGE MILL ROAD,

PALO ALTO, CA, 943041050

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 3 Drawing Page(s)

LINE COUNT: 1044

The present invention provides novel compositions and methods for AR lowering blood glucose levels, as well as manufacture processes for producing the compositions. Specifically, the present invention provides novel compositions that are extracts of the plant Prunella Linn and/or Rabdosis (Blume) Hasskarl containing enriched corosolic acid. Methods of isolating corosolic acid at high purity from these plants are also provided. These extracts and the purified corosolic acid can be used for lowering blood sugar levels and reducing accumulation of triglyeride in the treatment of diabetes, obesity and related conditions.

L19 ANSWER 84 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2003:159938 USPATFULL

Treatment of diabetes with thiazolidinedione and TITLE:

metformin

Smith, Stephen Alistair, Bramfield, UNITED KINGDOM INVENTOR(S):

PATENT ASSIGNEE(S): SmithKline Beecham plc (non-U.S. corporation)

NUMBER KIND DATE -----US 2003109553 A1 20030612 US 2003-340426 A1 20030110 (10) PATENT INFORMATION:

APPLICATION INFO.:

Continuation of Ser. No. US 2002-99161, filed on 13 Mar RELATED APPLN. INFO.:

2002, ABANDONED Continuation of Ser. No. US 2001-925394, filed on 9 Aug 2001, ABANDONED

Continuation of Ser. No. US 1999-446030, filed on 15 Dec 1999, ABANDONED A 371 of International Ser. No. WO

1998-EP3690, filed on 15 Jun 1998, UNKNOWN

NUMBER DATE -----

GB 1997-12857 19970618 PRIORITY INFORMATION: GB 1998-6706 19980327

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

GLAXOSMITHKLINE, Corporate Intellectual Property -LEGAL REPRESENTATIVE:

UW2220, P.O. Box 1539, King of Prussia, PA, 19406-0939

NUMBER OF CLAIMS: 22 EXEMPLARY CLAIM: 1 LINE COUNT: 484

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for the treatment and/or prophylaxis of diabetes mellitus, conditions associated with diabetes mellitus and certain complications thereof, in a mammal which method comprises administering an effective non-toxic and pharmaceutically acceptable amount of an insulin

sensitiser and a biguanide antihyperglycaemic agent, to a mammal in need thereof and a pharmaceutical composition comprising an insulin sensitiser and a biguanide antihyperglycaemic agent.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 85 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:251818 USPATFULL

Treatment of diabetes with thiazolidinedione and TITLE:

metformin

INVENTOR(S): Smith, Stephen Alistair, Bramfield, UNITED KINGDOM

PATENT ASSIGNEE(S): SmithKline Beecham plc (non-U.S. corporation)

NUMBER KIND DATE -----

US 2002137772 A1 20020926 US 2002-99161 A1 20020313 (10) PATENT INFORMATION: APPLICATION INFO.:

RELATED APPLN. INFO.: Continuation of Ser. No. US 2001-925394, filed on 9 Aug

2001, ABANDONED Continuation of Ser. No. US

1999-446030, filed on 15 Dec 1999, ABANDONED A 371 of International Ser. No. WO 1998-EP3690, filed on 15 Jun

1998, UNKNOWN

NUMBER DATE **---**----

GB 1997-12857 19970618 PRIORITY INFORMATION: GB 1998-6706

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

GLAXOSMITHKLINE, Corporate Intellectual Property -LEGAL REPRESENTATIVE:

UW2220, P.O. Box 1539, King of Prussia, PA, 19406-0939

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT: 485

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for the treatment and/or prophylaxis of diabetes mellitus, conditions associated with diabetes mellitus and certain complications thereof, in a mammal which method comprises administering an effective non-toxic and pharmaceutically acceptable amount of an insulin sensitiser and a biguanidine antihyperglycaemic agent, to a mammal in need thereof and a pharmaceutical composition comprising an insulin sensitiser and a biguanide antihyperglycaemic agent.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 86 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:8515 USPATFULL

TITLE: Treatment of diabetes with thiazolidinedione and

metformin

INVENTOR(S): Smith, Stephen Alistair, Bramfield, UNITED KINGDOM

NUMBER KIND DATE ------US 2002004515 A1 20020110 US 2001-925394 A1 20010809 (9) PATENT INFORMATION: APPLICATION INFO.:

RELATED APPLN. INFO.: Continuation of Ser. No. US 1999-446030, filed on 15 Dec 1999, ABANDONED A 371 of International Ser. No. WO

1998-EP3690, filed on 15 Jun 1998, UNKNOWN

NUMBER DATE -----

PRIORITY INFORMATION: GB 1997-12857 19970618 GB 1998-6706

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

GLAXOSMITHKLINE, Corporate Intellectual Property -LEGAL REPRESENTATIVE:

UW2220, P.O. Box 1539, King of Prussia, PA, 19406-0939

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 LINE COUNT: 484

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for the treatment and/or prophylaxis of diabetes mellitus, conditions associated with diabetes mellitus and certain complications thereof, in a mammal which method comprises administering an effective non-toxic and pharmaceutically acceptable amount of an insulin sensitiser and a biguanidine antihyperglycaemic agent, to a mammal in need thereof and a pharmaceutical composition comprising an insulin sensitiser and a biguanide antihyperglycaemic agent.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 87 OF 256 USPATFULL on STN

2002:54399 USPATFULL ACCESSION NUMBER:

TITLE: Preparation of aqueous clear solution dosage forms with

bile acids

Yoo, Seo Hong, Wyckoff, NJ, UNITED STATES INVENTOR(S):

NUMBER KIND DATE -----PATENT INFORMATION: US 2002031558 A1 20020314 US 2001-778154 A1 20010205 (9) APPLICATION INFO.:

Continuation-in-part of Ser. No. US 1999-357549, filed RELATED APPLN. INFO.:

on 20 Jul 1999, GRANTED, Pat. No. US 6251428

NUMBER DATE -----

US 1998-94069P 19980724 (60) PRIORITY INFORMATION:

US 2000-180268P 20000204 (60)

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: BAKER BOTTS L.L.P., 44TH FLOOR, 30 ROCKEFELLER PLAZA,

NEW YORK, NY, 10112-4498

NUMBER OF CLAIMS: 87 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 12 Drawing Page(s)

LINE COUNT: 2250

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Compositions for pharmaceutical and other uses comprising clear aqueous solutions of bile acids which do not form any detectable precipitates over selected ranges of pH values of the aqueous solution and methods of making such solutions. The compositions of the invention comprise water; a bile acid in the form of a bile acid, bile acid salt, or a bile acid conjugated with an amine by an amide linkage; and either or both an aqueous soluble starch conversion product and an aqueous soluble non-starch polysaccharide. The composition remains in solution without forming a precipitate over a range of pH values and, according to one embodiment, remains in solution for all pH values obtainable in an aqueous system. The composition, according to some embodiments, may further contain a pharmaceutical compound in a pharmaceutically effective amount. Non-limiting examples of pharmaceutical compounds include insulin, heparin, bismuth compounds, amantadine and rimantadine.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 88 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2001:228693 HCAPLUS

DOCUMENT NUMBER: 134:256878

TITLE: Pharmaceuticals containing nateglinide or repaglinide for treating diabetes or conditions assocd. with

diabetes

INVENTOR(S): Gatlin, Marjorie Regan; Pongowski, Michele; Mannion,

Richard Owen; Karnachi, Anees Abdulquadar; Guitard,

Christiane; Allison, Malcolm

PATENT ASSIGNEE(S): Novartis A.-G., Switz.; Novartis-Erfindungen

Verwaltungsgesellschaft m.b.H.

SOURCE: PCT Int. Appl., 60 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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KIND DATE
                                       APPLICATION NO. DATE
    PATENT NO.
                                        -----
                   _____
                                       WO 2000-EP9074 20000915
                          20010329
    WO 2001021159
                   A2
    WO 2001021159
                    A3
                          20011227
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
            HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
            LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
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            YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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            CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                      FR 2000-11782
                     A1 20010323
    FR 2798592
                                        BR 2000-14525
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    EP 1212077
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                                        EP 2000-969260
                                                        20000915
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL
                                        BE 2000-585
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                    A5
                        20020702
    BE 1013726
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    JP 2003509457
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    US 6559188
                                        US 2000-663264
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                          20010402
                                        FI 2001-683
                                                        20010402
    FI 2001000683
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                                        NO 2002-1197
    NO 2002001197
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                          20020516
                                                        20020311
                    A1 20030828
                                        US 2003-345908
                                                        20030116
    US 2003162816
PRIORITY APPLN. INFO.:
                                     US 1999-242911P P 19990917
                                     US 1999-398364 A 19990917
                                     US 2000-240918P P 20000309
                                     US 2000-304196P P 20000407
                                     US 2000-545480 A 20000407
                                                     A 20000826
                                      GB 2000-21055
                                     US 1999-240911P P 19990917
                                      US 2000-521737 A 20000309
                                      US 2000-663264 A1 20000915
                                      WO 2000-EP9074 W 20000915
```

The invention relates to a combination, such as a combined prepn. or pharmaceutical compn., resp., which comprises nateglinide or repaglinide and at 1 other antidiabetic compd. selected from the group consisting of thiazolidinedione derivs. (glitazones), sulfonylurea derivs. and metformin for simultaneous, sep. or sequential use in the prevention, delay of progression or treatment of the diseases. The compn. is esp. useful for the treatment of type 2 diabetes and diseases. Thus, tablet contained nateglinide 12.960, lactose 30.564, microcryst. cellulose 15.336, povidone 2.592, croscarmellose sodium 3.974, colloidal SiO2 1.382, magnesium stearate 1.231, and coating with Opadry yellow 1.944 kg., and water qs.

L19 ANSWER 89 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:112964 USPATFULL

TITLE: COMPOSITIONS CONTAINING HYPOGLYCEMICALLY ACTIVE

STILBENOIDS

INVENTOR(S): Hopp, David C., Mill Creek, WA, UNITED STATES

Inman, Wayne D., Belmont, CA, UNITED STATES

NUMBER DATE

PRIORITY INFORMATION: US 2000-225800P 20000816 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: STERNE, KESSLER, GOLDSTEIN & FOX PLLC, 1100 NEW YORK

AVENUE, N.W., SUITE 600, WASHINGTON, DC, 20005-3934

NUMBER OF CLAIMS: 16 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 6 Drawing Page(s)

LINE COUNT: 2015

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The use of isolated or purified stilbenoid compounds including longistyline A-2-carboxylic acid as hypoglycemic agents or to lower serum glucose levels is described. The invention also relates to the use of such stibenoid compounds in combination with other hypoglycemic

agents.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 90 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:112958 USPATFULL

TITLE: Compositions containing hypoglycemically active

stilbenoids

INVENTOR(S): Inman, Wayne D., Belmont, CA, UNITED STATES

Hopp, David C., Mill Creek, WA, UNITED STATES

NUMBER DATE

PRIORITY INFORMATION: US 2000-225665P 20000816 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: STERNE, KESSLER, GOLDSTEIN & FOX PLLC, 1100 NEW YORK

AVENUE, N.W., SUITE 600, WASHINGTON, DC, 20005-3934

NUMBER OF CLAIMS: 16 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 6 Drawing Page(s)

LINE COUNT: 2013

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The use of isolated or purified stilbenoid compounds including longistyline A-2-carboxylic acid as hypoglycemic agents or to lower serum glucose levels is described. The invention also relates to the use of such stibenoid compounds in combination with other hypoglycemic

agents.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 91 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1999:549136 HCAPLUS

DOCUMENT NUMBER: 131:161654

Orally administrable immediate-release and TITLE:

prolonged-release galenic form comprising an

absorption-promoting agent

Saslawski, Olivier; Giet, Philippe; Michel, Dominique; INVENTOR (S):

Hulot, Thierry

Merck Patent G.m.b.H., Germany PATENT ASSIGNEE(S):

SOURCE: PCT Int. Appl., 65 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                   KIND DATE
                                       APPLICATION NO. DATE
                   A1 19990826 WO 1999-EP994 19990216
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    WO 9942086
        W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
           DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP,
            KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,
            MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
            TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU,
            TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
            FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
            CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    FR 2775188
                    A1
                          19990827
                                       FR 1998-2143
                                                        19980223
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                          20010309
    CA 2321267
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                          19990826
                                        CA 1999-2321267 19990216
    AU 9931408
                    A1
                          19990906
                                        AU 1999-31408
                                                        19990216
    AU 750785
                    B2
                          20020725
                          20001024
                                       BR 1999-8121
                                                        19990216
    BR 9908121
                   Α
                        20001206
                                       EP 1999-913165 19990216
    EP 1056445
                    A1
           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE,
            SI, LT, LV, FI, RO
    JP 2002503686
                    T2
                          20020205
                                        JP 2000-532103
                                                        19990216
    ZA 9901408
                          19990823
                                        ZA 1999-1408
                                                        19990222
                     Α
    NO 2000004190
                          20001020
                                        NO 2000-4190
                                                        20000822
                     Α
    US 6426087
                     B1
                          20020730
                                        US 2000-622663
                                                       20000822
                         20030204
                                        US 2002-100084 20020319
    US 6514524
                    B1
PRIORITY APPLN. INFO.:
                                     FR .1998-2143 A 19980223
                                     WO 1999-EP994
                                                     W 19990216
                                     US 2000-622633 A1 20000822
```

OTHER SOURCE(S): MARPAT 131:161654

The present invention relates to an orally administrable galenic form allowing improved absorption by the transmembrane or paracellular route in the gastrointestinal tract of active ingredients which are hydrophilic or ionizable in physiol. media, comprising at least one such active ingredient, an absorption-promoting agent having an HLB >8, the the absorption-promoting agent consisting of one or more lipid substances chosen from: polysorbates; polyoxyethylene ethers; esters of polyoxyethylene and fatty acids; fatty acids; fatty alcs.; bile acids and their salts with pharmaceutically acceptable cations; esters of C1-C6 alkanol with fatty acids; esters of polyol with fatty acids, the polyol comprising from 2 to 6 hydroxyl functional groups; and polyglycolyzed glycerides; in combination with one or more pharmaceutically acceptable excipients, the pharmaceutical forms comprising captopril being excluded. A controlled-release tablet contained (1) cores contg. calcium acamprosate 50, Gelucire 44/14 10, Compritol 10, microcryst. cellulose 19, Povidone 10, and Mg stearate 1 % and (2) a filmcoating compn. contg. HPMC 64, PEG-4000 15, and talc 21 %.

REFERENCE COUNT: THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT L19 ANSWER 92 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2003:472994 HCAPLUS

DOCUMENT NUMBER: 139:41844

TITLE: Reverse micellar delivery system for controlled

transport and enhanced drug absorption

INVENTOR(S): MacGregor, Alexander

PATENT ASSIGNEE(S): Can

SOURCE: U.S. Pat. Appl. Publ., 16 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND DAT	E	APPLICATIO	ON NO.	DATE	
US 2003113366	A1 200	30619	US 2001-24	1325	20011214	
WO 2003051333	A1 200	30626	WO 2002-CA	1918	20021213	
W: AE, AG,	AL, AM, AT	, AU, AZ,	BA, BB, BG,	BR, BY,	BZ, CA,	CH, CN,
CO, CR,	CU, CZ, DE	E, DK, DM,	DZ, EC, EE,	ES, FI,	GB, GD,	GE, GH,
GM, HR,	HU, ID, II	, IN, IS,	JP, KE, KG,	KP, KR,	KZ, LC,	LK, LR,
LS, LT,	LU, LV, MA	A, MD, MG,	MK, MN, MW,	MX, MZ,	NO, NZ,	OM, PH,
PL, PT,	RO, RU, SO	C, SD, SE,	SG, SK, SL,	TJ, TM,	TN, TR,	TT, TZ,
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PT, SE,	SI, SK, TF	R, BF, BJ,	CF, CG, CI,	CM, GA,	GN, GQ,	GW, ML,
MR. NE.	SN, TD, TO	}				

PRIORITY APPLN. INFO.: US 2001-24325 A 20011214

The present invention provides a reverse-micellar delivery system for enhanced absorption of an agent of interest across biol. membranes such as the gastro-intestinal tract of mammals. The reverse-micelles comprise at least one ionic amphipathic compd., and at least one polar active agent ionizable in aq. or physiol. media. The delivery system facilitates transportation of the agent across the gastro-intestinal tract or other membranes and enhances the in-vivo release and availability of the agent(s) of interest within a fluid environment. An extended release tablet contained metformin-HCl 69, cetyl alc. 18, Na lauryl sulfate 10, Et cellulose 2, and Mg stearate 1%/tablet.

L19 ANSWER 93 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2001:163422 HCAPLUS

DOCUMENT NUMBER: 134:212730

TITLE: Controlled-release lipoic acid
INVENTOR(S): Byrd, Edward A.; Janjikhel, Rajiv
PATENT ASSIGNEE(S): Medical Research Institute, USA

SOURCE: U.S., 14 pp., Cont.-in-part of U.S. Ser. No. 112,623,

abandoned.
CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6197340	B1	20010306	US 1999-288245	19990408
US 6191162	B1	20010220	US 1999-288253	19990408
CA 2332790	AA	19991202	CA 1999-2332790	19990519
WO 9961004	A1	19991202	WO 1999-US11178	19990519
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DE, DK,	EE, ES	, FI, GB, GD,	GE, GH, GM, HR, HU	, ID, IL, IN, IS,

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             MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
             TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD,
             RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
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             CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     AU 9940903
                      A1
                           19991213
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     EP 1082107
                      Α1
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                                          EP 1999-924394
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         R: DE, ES, FR, GB, IT
     JP 2002516270
                   T2 20020604
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     US 2001028896
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                                          US 2001-755890
                                                           20010105
     US 6572888
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                           20030603
     US 2003039690
                     A1 20030227
                                          US 2002-226646
                                                           20020823
PRIORITY APPLN. INFO.:
                                       US 1998-87203P P 19980528
                                       US 1998-112623 B2 19980709
                                       US 1998-102605P P 19981001
                                       US 1999-288245 A 19990408
                                       WO 1999-US11178 W 19990519
                                       US 2001-755890
                                                       A2 20010105
AΒ
     A controlled release formulation of lipoic acid is disclosed.
                                                                   The lipoic
     acid is combined with excipient materials in such a way that those
     materials protect the lipoic acid from chem. degrdn. in the
     gastrointestinal tract and provide for gradual release of the lipoic acid.
     These combined features make it possible to use lipoic acid to reduce
     serum glucose levels and maintain those levels over time thereby obtaining
     a range of desired results. A sustained-release tablet
     contained racemic .alpha.-lipoic acid coated particles 81, Methocel K100
     10, microcryst. cellulose 5, stearic acid 3, micronized silica
     0.5, and magnesium stearate 0.5%. Efficacy of the formulation in lowering
     blood glucose level of patients is reported.
REFERENCE COUNT:
                        115
                              THERE ARE 115 CITED REFERENCES AVAILABLE FOR
                              THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE
                              FORMAT
L19 ANSWER 94 OF 256 USPATFULL on STN
ACCESSION NUMBER:
                       2003:24185 USPATFULL
TITLE:
                       Combination therapy for type II diabetes or Syndrome X
                       Gwynne, John Thomas, Doylestown, PA, UNITED STATES
INVENTOR(S):
                       Vitou, Philippe John Robert, Paris, FRANCE
                       Randazzo, Bruce Paul, Rydal, PA, UNITED STATES
PATENT ASSIGNEE(S):
                       Wyeth, Madison, NJ (U.S. corporation)
                            NUMBER
                                         KIND
                                               DATE
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PATENT INFORMATION:
                       US 2003018028
                                          A1
                                               20030123
APPLICATION INFO.:
                       US 2002-163707
                                          A1
                                               20020606 (10)
                                           DATE
                             NUMBER
PRIORITY INFORMATION:
                       US 2001-296502P
                                          20010607 (60)
DOCUMENT TYPE:
                       Utility
FILE SEGMENT:
                       APPLICATION
LEGAL REPRESENTATIVE:
                       Arnold S. Milowsky, 5 Giralda Farms, Madison, NJ, 07940
NUMBER OF CLAIMS:
                       23
EXEMPLARY CLAIM:
LINE COUNT:
                       1108
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention provides methods of using a pharmacological combination
AB
       of a biguanide agents, such as metformin, and one or more
      PTPase inhibiting agents and, optionally, one or more sulfonlylurea
       agents, including glyburide, glyburide, glipizide, glimepiride,
       chlorpropamide, tolbutamide, or tolazamide, for treatment in a mammal of
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Syndrome X, type II diabetes or metabolic disorders mediated by insulin

resistance or hyperglycemia. Further included in this invention is a method of modulating blood glucose levels in a mammal utilizing the combination of one or more PTPase inhibiting agents and one or more sulfonlylurea agents.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 95 OF 256 USPATFULL on STN

1999:160079 USPATFULL ACCESSION NUMBER:

Glycogen phosphorylase inhibitors TITLE:

Hulin, Bernard, Essex, CT, United States INVENTOR(S):

Sarges, Reinhard, Mystic, CT, United States

Pfizer Inc, New York, NY, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE

-----PATENT INFORMATION: US 5998463 19991207 APPLICATION INFO.: US 1999-251141 19990216

19990216 (9)

NUMBER DATE ______

PRIORITY INFORMATION: US 1998-76132P 19980227 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Richter, Johann ASSISTANT EXAMINER: Keating, Dominic

LEGAL REPRESENTATIVE: Richardson, Peter C., Benson, Gregg C., Gammill, Martha

Α.

NUMBER OF CLAIMS: 23 EXEMPLARY CLAIM: 1 LINE COUNT: 1835

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

This invention relates to certain 5-acyl-2-oxo-indole-3-carboxamides useful as inhibitors of glycogen phosphorylase, methods of treating glycogen phosphorylase dependent diseases or conditions with such compounds and pharmaceutical compositions comprising such compounds. This invention also relates to pharmaceutical compositions comprising those 5-acyl-2-oxo-indole-3-carboxamides in combination with antidiabetes agents and methods of treating glycogen phosphorylase dependent diseases or conditions with such compositions.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 96 OF 256 USPATFULL on STN

2003:79163 USPATFULL ACCESSION NUMBER:

Combination therapy comprising glucose reabsorption TITLE:

inhibitors and retinoid-X receptor modulators

Bussolari, Jacqueline C., Skillman, NJ, UNITED STATES INVENTOR(S):

Chen, Xiaoli, Belle Mead, NJ, UNITED STATES Conway, Bruce R., Doylestown, PA, UNITED STATES Demarest, Keith T., Flemington, NJ, UNITED STATES Ross, Hamish N.M., Far Hills, NJ, UNITED STATES

Severino, Rafael, Madrid, SPAIN

KIND DATE NUMBER -----

US 2003055091 A1 20030320 US 2002-115725 A1 20020403 (10) PATENT INFORMATION: APPLICATION INFO.:

> NUMBER DATE _____

US 2001-281479P 20010404 (60) PRIORITY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

AUDLEY A. CIAMPORCERO JR., JOHNSON & JOHNSON, ONE LEGAL REPRESENTATIVE:

JOHNSON & JOHNSON PLAZA, NEW BRUNSWICK, NJ, 08933-7003

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 LINE COUNT: 2308

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Combination therapy comprising RXR modulators and glucose reabsorption

inhibitors useful for the treatment of diabetes and Syndrome X are

disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 97 OF 256 USPATFULL on STN

2003:65429 USPATFULL ACCESSION NUMBER:

TITLE:

Combination therapy comprising glucose reabsorption

inhibitors and PPAR modulators

INVENTOR(S):

Bussolari, Jacqueline C., Skillman, NJ, UNITED STATES

Chen, Xiaoli, Belle Mead, NJ, UNITED STATES Conway, Bruce R., Doylestown, PA, UNITED STATES Demarest, Keith T., Flemington, NJ, UNITED STATES Ross, Hamish N.M., Far Hills, NJ, UNITED STATES

Severino, Rafael, Madrid, SPAIN

KIND DATE NUMBER -----

PATENT INFORMATION: APPLICATION INFO.:

US 2003045553 A1 20030306 US 2002-115827 A1 20020403 (10)

NUMBER DATE ______

PRIORITY INFORMATION:

US 2001-281429P 20010404 (60)

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: AUDLEY A. CIAMPORCERO JR., JOHNSON & JOHNSON, ONE

JOHNSON & JOHNSON PLAZA, NEW BRUNSWICK, NJ, 08933-7003

NUMBER OF CLAIMS: 67 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 7 Drawing Page(s)

LINE COUNT: 2106

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Combination therapy comprising PPAR modulators and glucose reabsorption

inhibitors useful for the treatment of diabetes and Syndrome X are

disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L19 ANSWER 98 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1998:804171 HCAPLUS

DOCUMENT NUMBER:

130:57204

TITLE:

Gastric-retentive oral drug dosage forms for controlled release of highly soluble drugs

INVENTOR(S):

Shell, John W.; Louie-Helm, Jenny

PATENT ASSIGNEE(S):

Depomed, Inc., USA PCT Int. Appl., 31 pp.

SOURCE: CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

KIND DATE APPLICATION NO. DATE PATENT NO. ----------WO 9855107 A1 19981210 WO 1998-US11302 19980605

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W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
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            KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
            NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
            UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
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                          19981221
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                                                         19980605
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    EP 998271
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    US 2001018070
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                           20010830
                           20020122
    US 6340475
                      B2
                                       US 1997-870509 A2 19970606
PRIORITY APPLN. INFO.:
                                       WO 1998-US11302 W 19980605
    Drugs that are freely or highly sol. in water are formulated as unit
AΒ
    dosage forms by incorporating them into polymeric matrixes comprised of
    high mol. wt. hydrophilic polymers that swell upon imbibition of water.
    The dosage form can be a single compressed tablets, or two or
    three compressed tablets retained in a single gelatin
    capsule. The oral formulation is designed for gastric retention
    and controlled delivery of an incorporated drug into the gastric cavity,
    and thus administered, the drug is released from the matrix into the
    gastric fluid by soln. diffusion. The swollen polymeric matrix, having
    achieved sufficient size, remains in the gastric cavity for several hours
     if administered while the patient is in the fed mode, and remains intact
     long enough for substantially all of the drug to be released before
     substantial erosion of the matrix occurs. The swelling matrix lowers the
     accessibility of the gastric fluid to the drug and thereby limits the drug
    release rate. This process, together with diffusion retardation by
     selection of specific polymers, polymer mol. wts., and other variables,
     results in a sustained and controlled delivery rate of the drug to the
     gastric environment. Controlled-release behavior of metformin
     -HCl from a polyethylene oxide matrix was demonstrated.
                              THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                        5
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L19 ANSWER 99 OF 256 HCAPLUS COPYRIGHT 2003 ACS on STN
                        2003:334870 HCAPLUS
ACCESSION NUMBER:
                        138:343894
DOCUMENT NUMBER:
                        Formulation of an erodible, gastric retentive oral
TITLE:
                        dosage form using in vitro disintegration test data
                        Louie-helm, Jenny; Berner, Bret
INVENTOR(S):
                        Depomed, Inc., USA
PATENT ASSIGNEE(S):
                        PCT Int. Appl., 55 pp.
SOURCE:
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                   KIND DATE
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                                        WO 2002-US34298 20021025
                     A1 20030501
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        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
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WO 2003035029 A1 20030501 WO 2002-US34298 20021025
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,

NE, SN, TD, TG

US 2003091630 A1 20030515 US 2001-14750 20011025 PRIORITY APPLN. INFO.: US 2001-14750 A 20011025

AB Erodible, gastric-retentive dosage forms are provided that are formulated using the in vitro drug release profile obtained with USP disintegration test equipment rather the USP Dissoln. App. The invention is premised on the discovery that the USP disintegration test and modified versions thereof are far more predictive of the in vivo release profile for a controlled release dosage form than is the std. USP disintegration test, particularly controlled release dosage forms of the swellable, erodible type. The dosage forms generally comprise particles of a biocompatible, hydrophilic polymer having the active agent incorporated therein, wherein the particles are optionally but preferably compacted into a tablet or loaded into a capsule. The dosage forms can be used to deliver water-insol. or sparingly sol. drugs as well as water-sol. drugs, providing that the latter are coated with a protective coating or contained in a protective vesicle. Tablet

coating or contained in a protective vesicle. Tablet
contained BaSO4 21.35, Polyox N-60K 20.02, and Polyox N-80 58.13%.

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L19 ANSWER 100 OF 256 USPATFULL on STN

ACCESSION NUMBER: 2002:185267 USPATFULL

TITLE: Dietetic food composition and dietetic method using

such composition

INVENTOR(S): Zohoungbogbo, Mathias C., Torino, ITALY

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-333097, filed on 15 Jun 1999, PATENTED Continuation-in-part of Ser.

No. US 1999-225819, filed on 5 Jan 1999, ABANDONED

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: SOFER & HAROUN, L.L.P., Suite 1921, 342 Madison Avenue,

New York, NY, 10173

NUMBER OF CLAIMS: 25 EXEMPLARY CLAIM: 1 LINE COUNT: 709

PATENT INFORMATION: APPLICATION INFO.:

PRIORITY INFORMATION:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Food composition in the form of a flour comprising at least 50% of protein, less than 15% of carbohydrates and 35 to 50% of plant fibers; preferably the carbohydrate content is less than 10%, advantageously less than 5%; this composition may be used as a substitute for wheat flour in the preparation of foods such as pasta, bread, bread sticks, bakery products and pastries and constitutes the basis of a method for improving the appearance of a person by achieving a loss of weight which is beneficial from the aesthetic point of view.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

FISCAL YEAR 2004

Dates	Oct-01 - Oct-18-2003	Oct-19 - Nov-01-2003	Nov-02 - Nov-15-2003	Nov-16 - Nov-29-2003	Nov-30 - Dec-13-2003	(End of 1st Quarter)	Dec-14 - Dec-27-2003	Dec-28 - Jan-10-2004	Jan-11 - Jan-24-2004	Jan-25 - Feb-07-2004	Feb-08 - Feb-21-2004	Feb-22 - Mar-06-2004	Mar-07 - Mar-20-2004	(End of 2nd Quarter)	Mar-21 - Apr-03-2004	Apr-04 - Apr-17-2004	Apr-18 - May-01-2004	May-02 - May-15-2004	May-16 - May-29-2004	May-30 - Jun-12-2004	Jun-13 - Jun-26-2004	(End of 3rd Quarter)	Jun-27 - Jul-10-2004	Jul-11 - Jul-24-2004	Jul-25 - Aug-07-2004	·	Aug-22 - Sep-04-2004	Sep-05 - Sep-18-2004	Sep-19 - Sep-30-2004	(End of 4th Quarter - EOY)
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1	1	"6303146" .pn.	USPAT;	2003/09/12 11:03
*		, 0303110 .pm.	US-PGPUB	2003,03,12 11.03
2	1222	metformin metomin miformidimethylbiquanide	USPAT;	2003/09/12 11:05
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3	1029	glipizide glucotrol glydiazinamide	USPAT;	2003/09/12 11:07
		glupizide glupitel glucozide glynase	US-PGPUB	
		minidab glican glidiab digrin dipazide		
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5	648	((metformin metomin	USPAT;	2003/09/12 11:14
	""	miformidimethylbiquanide glucomine	US-PGPUB	2000,03,12 11.11
		glucophage glufor gluformin diformin		
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		and (glipizide glucotrol glydiazinamide		
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		and (coat or coating or (magnesium adj		
		stearate) or pyrrolidone or cellulose or		
		croscarmellose or (single adj dosage) or		
6	490	tablet or capsule) (((metformin metomin	USPAT;	2003/09/12 11:10
0	490	miformidimethylbiguanide glucomine	US-PGPUB	2003/09/12 11:10
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		and (glipizide glucotrol glydiazinamide		
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7	17	((((metformin metomin	USPAT;	2003/09/12 11:10
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		stearate) or pyrrolidone or cellulose or croscarmellose or (single adj dosage) or		
		tablet or capsule)) and (magnesium adj		
1		stearate)) not combination		
8	23	(((metformin metomin	USPAT;	2003/09/12 11:10
	_	miformidimethylbiguanide glucomine	US-PGPUB	
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9	134	((metformin metomin miformidimethylbiguanide glucomine glucophage glufor gluformin diformin dimefor geamet glucamino glucofago glucoform glucomin glucomine gluconil) and (glipizide glucotrol glydiazinamide glupizide glupitel glucozide glynase minidab glican glidiab digrin dipazide)) and ((magnesium adj stearate) and pyrrolidone or cellulose and	USPAT; US-PGPUB	2003/09/12 11:14
	75	croscarmellose) (((metformin metomin miformidimethylbiguanide glucomine glucophage glufor gluformin diformin dimefor geamet glucamino glucofago glucoform glucomin glucomine gluconil) and (glipizide glucotrol glydiazinamide glupizide glupitel glucozide glynase minidab glican glidiab digrin dipazide)) and ((magnesium adj stearate) and pyrrolidone or cellulose and	USPAT; US-PGPUB	2003/09/12 11:15
11	17	croscarmellose)) and (coating or coat) (((metformin metomin miformidimethylbiguanide glucomine glucophage glufor gluformin diformin dimefor geamet glucamino glucofago glucoform glucomin glucomine gluconil) and (glipizide glucotrol glydiazinamide glupizide glupitel glucozide glynase minidab glican glidiab digrin dipazide)) and ((magnesium adj stearate) and pyrrollone or cellulose and	USPAT; US-PGPUB	2003/09/12 11:15
12	76	miformidimethylbiguanide glucomine glucophage glufor gluformin diformin dimefor geamet glucamino glucofago glucoform glucomin glucomine gluconil) and (glipizide glucotrol glydiazinamide glupizide glupitel glucozide glynase minidab glican glidiab digrin dipazide)) and ((magnesium adj stearate) and pyrrolidone or cellulose and	USPAT; US-PGPUB	2003/09/12 12:25
13	281	croscarmellose)) and (opadry or coating) chen and metformin	USPAT; US-PGPUB; DERWENT	2003/09/12 12:26
14	233	(chen and metformin) and glipizide	USPAT; US-PGPUB; DERWENT	2003/09/12 12:26
15	4	((chen and metformin) and glipizide) and andrx	USPAT; US-PGPUB; DERWENT	2003/09/12 12:26



Creation date: 03-04-2004

Indexing Officer: AAHMED4 - ASHA AHMED

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1	IDS	4
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